



# NCAN

National Center for Adaptive Neurotechnologies

## Adaptive Neurotechnologies

*Rhythms Reflexes Rehabilitation*

Jonathan R. Wolpaw

National Center for Adaptive Neurotechnologies

Wadsworth Center, New York State Dept of Health



# Two Recent Advances

- Recognition that the CNS changes throughout life.  
*Activity-dependent plasticity* is occurring everywhere.
- Hardware & software for complex real-time interactions with the CNS that initiate & guide plasticity.

*Unprecedented opportunities for adaptive interactions:*  
*New insights*  
*New therapies*



# Adaptive Neurotechnologies

- The CNS interacts with the outside world & the body through peripheral nerves, sensory receptors, & muscles.
- Adaptive neurotechnologies *replace, restore, enhance, supplement, or improve* these natural interactions.
- These technologies typically adapt their behavior to the CNS & often induce adaptive plasticity in the CNS.



# Adaptive Neurotechnologies

- BCIs for restoring communication and control
- BCIs for stroke rehabilitation
- ECoG for pre-surgical mapping of cortical function
- Reflex conditioning for rehabilitation after spinal cord injury: Targeted Neuroplasticity



# What is a Brain-Computer Interface?

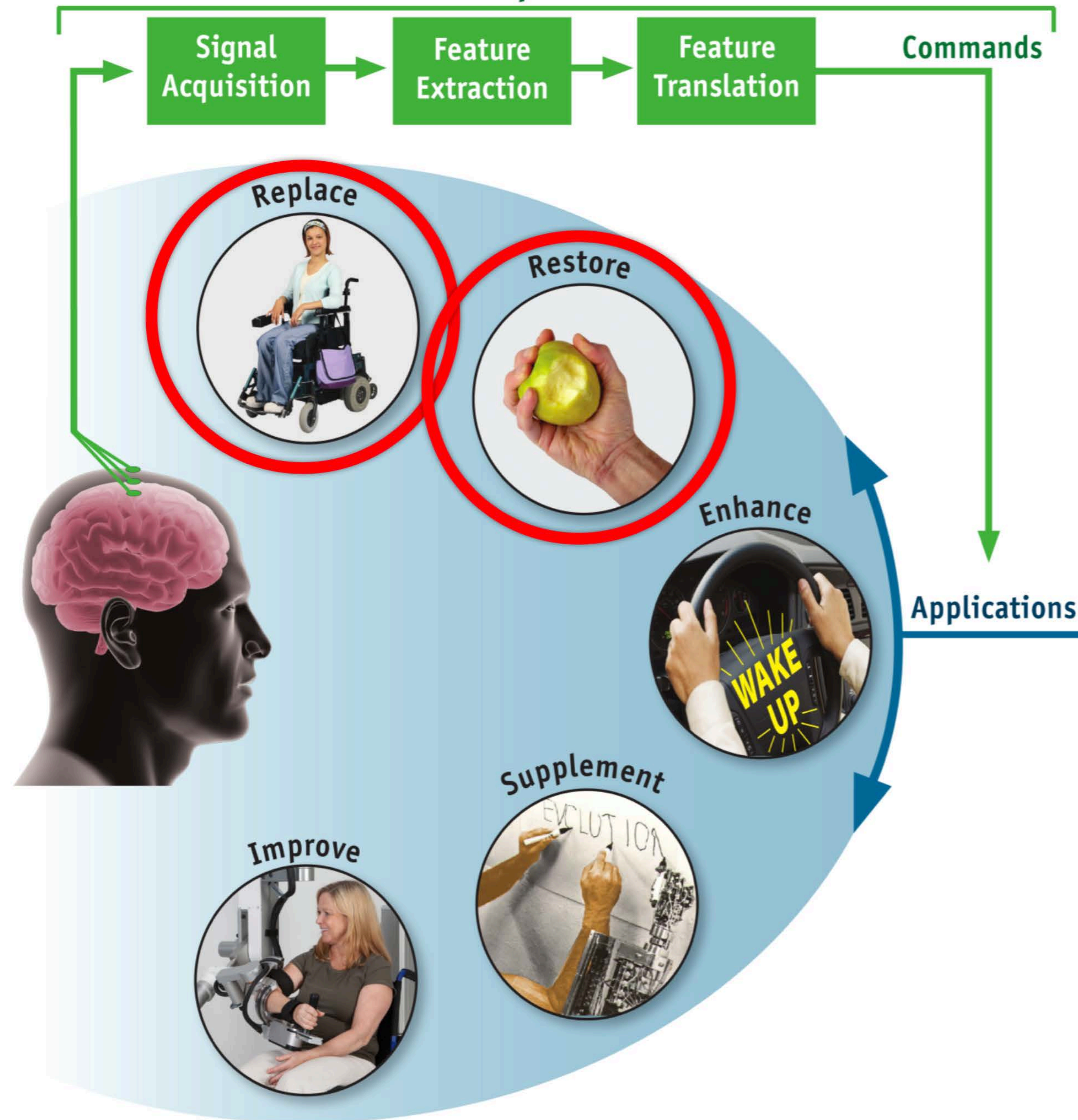
The CNS produces outputs that act on the world or the body. All its natural outputs are neuromuscular or hormonal.

*A BCI measures CNS activity and converts it into **artificial** output that **replaces, restores, enhances, supplements, or improves** natural output, and thereby changes the ongoing interactions between the CNS and its environment.*





# Brain-Computer Interface



**NCAN.**  
National Center for Adaptive Neurotechnology

Wolpaw & Wolpaw, Oxford Univ Press, 2012



**Department  
of Health**

Wadsworth  
Center



# Email with the Wadsworth BCI Home System

Wadsworth Email Client Send Email Mode

Send Email

Subject (Alt + u): Brain Computer Interface(BCI)

To (Alt + t): bci@wadsworth.org Cc (Alt + p):

hel

1 hello

2 help

3 held

4 helped

5 hell

6 helping

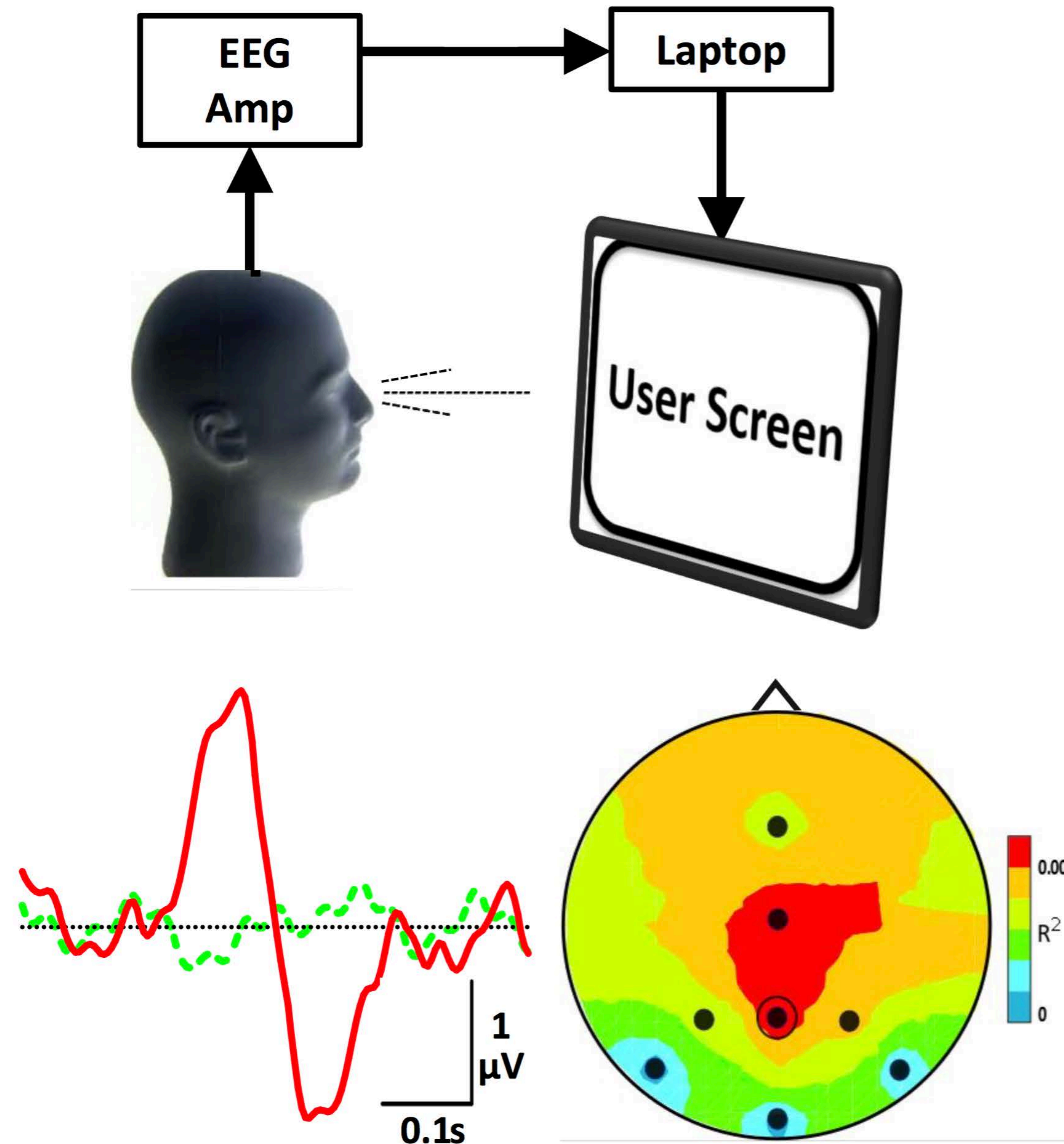
7 helps

A	B	C	D	E	F	G	H
I	J	K	L	M	N	O	P
Q	R	S	T	U	V	W	X
Y	Z	Sp	1	2	3	4	5
6	7	8	9	0	Prd	Ret	Bs
?	@	;	\	Help	+	-	Alt
Ctrl	=	Del	Home	UpAw	End	PgUp	Shft
Save	'	Load	LfAw	DnAw	RtAw	PgDn	Pause
Caps	Inbox	Tab	Compose	Esc	Send	!	Sleep

Alt+Shift+1	Alt+Shift+2	Alt+Shift+3	Alt+Shift+4	Alt+Shift+5	Alt+Shift+6
notepad.exe	emailclient.exe	Etriloquist5.exe	EMPTY	EMPTY	EMPTY
Focus Manager v. 1.0.3.96					



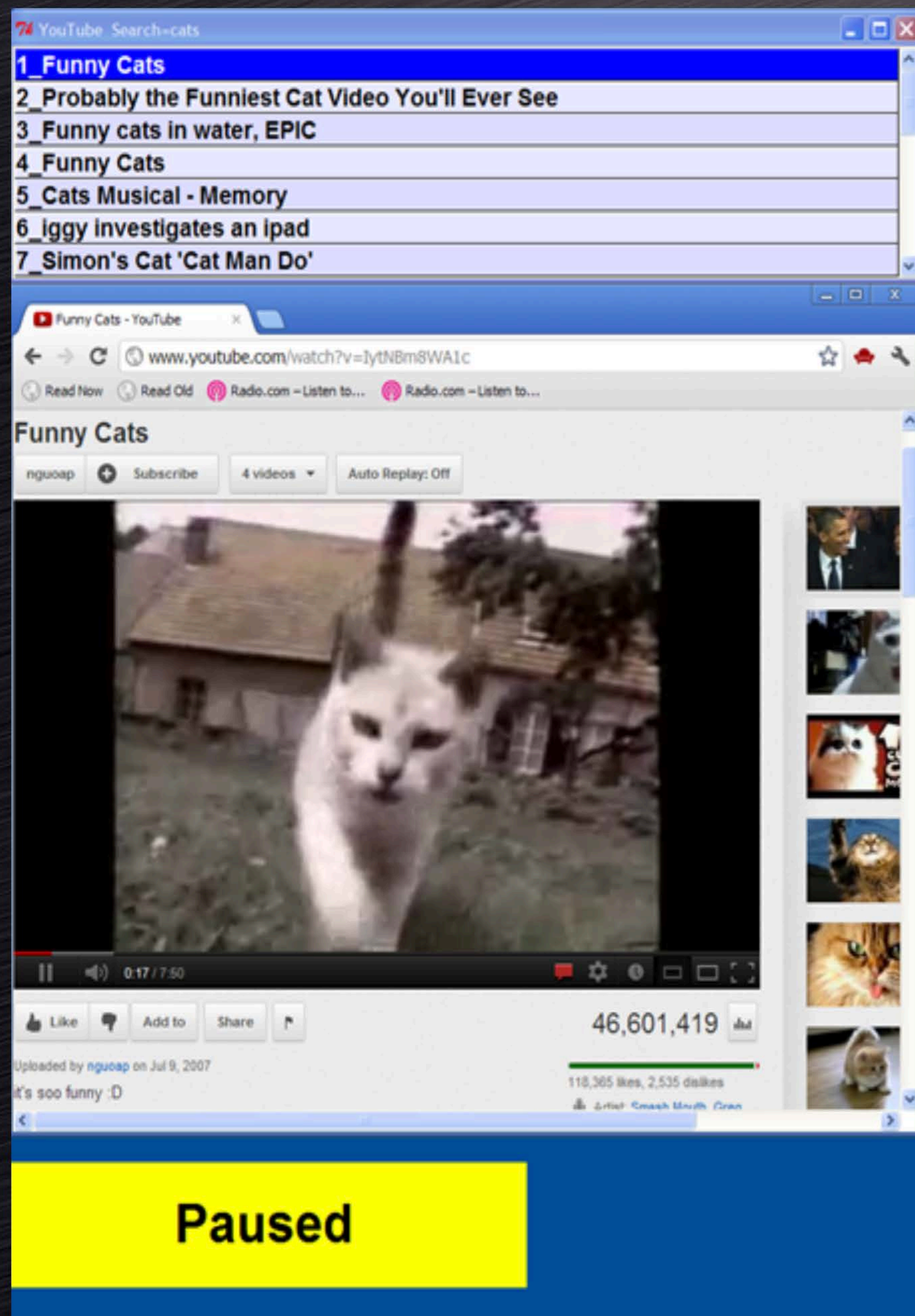
# P300 Evoked Potential



Donchin et al. 1988,2000



# BCI Home System Applications



A	B	C	D	E	F	G	H
I	J	K	L	M	N	O	P
Q	R	S	T	U	V	W	X
Y	Z	1	2	3	4	5	6
7	8	9	0	Sp	_	.	,
?	!	'	-	;	@	/	Enter
Bs	Del	Clear	Caps	PgUp	Home	↑	End
Esc	Ctrl	Alt	Shift	PgDn	←	↓	→
Vol+	Vol-	Help	Mute	RunNxt	Privacy	Pause	Menu



# Training Users and Their Caregivers





Asked to describe BCI impact on his life, the first user wrote:

▼ Subject: Re: [redacted]  
From: [redacted]  
Date: [redacted]  
To: Jonathan R. Wolpaw [redacted]  
Cc: [redacted]

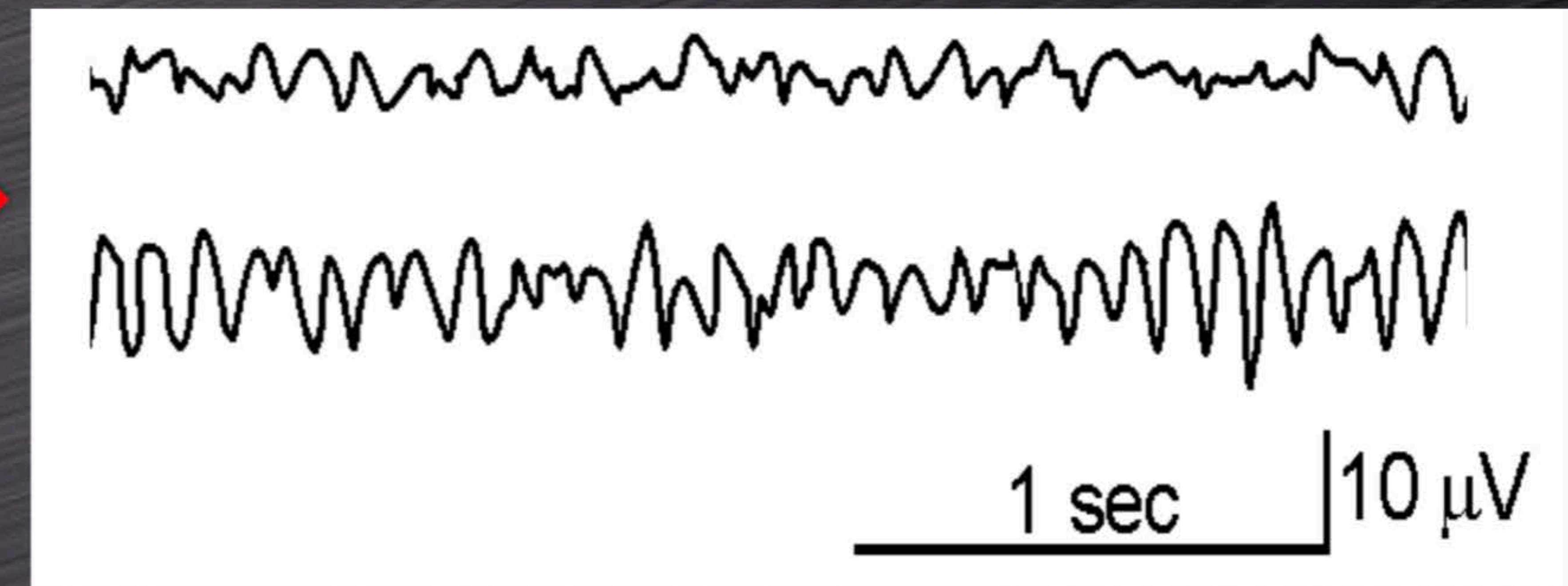
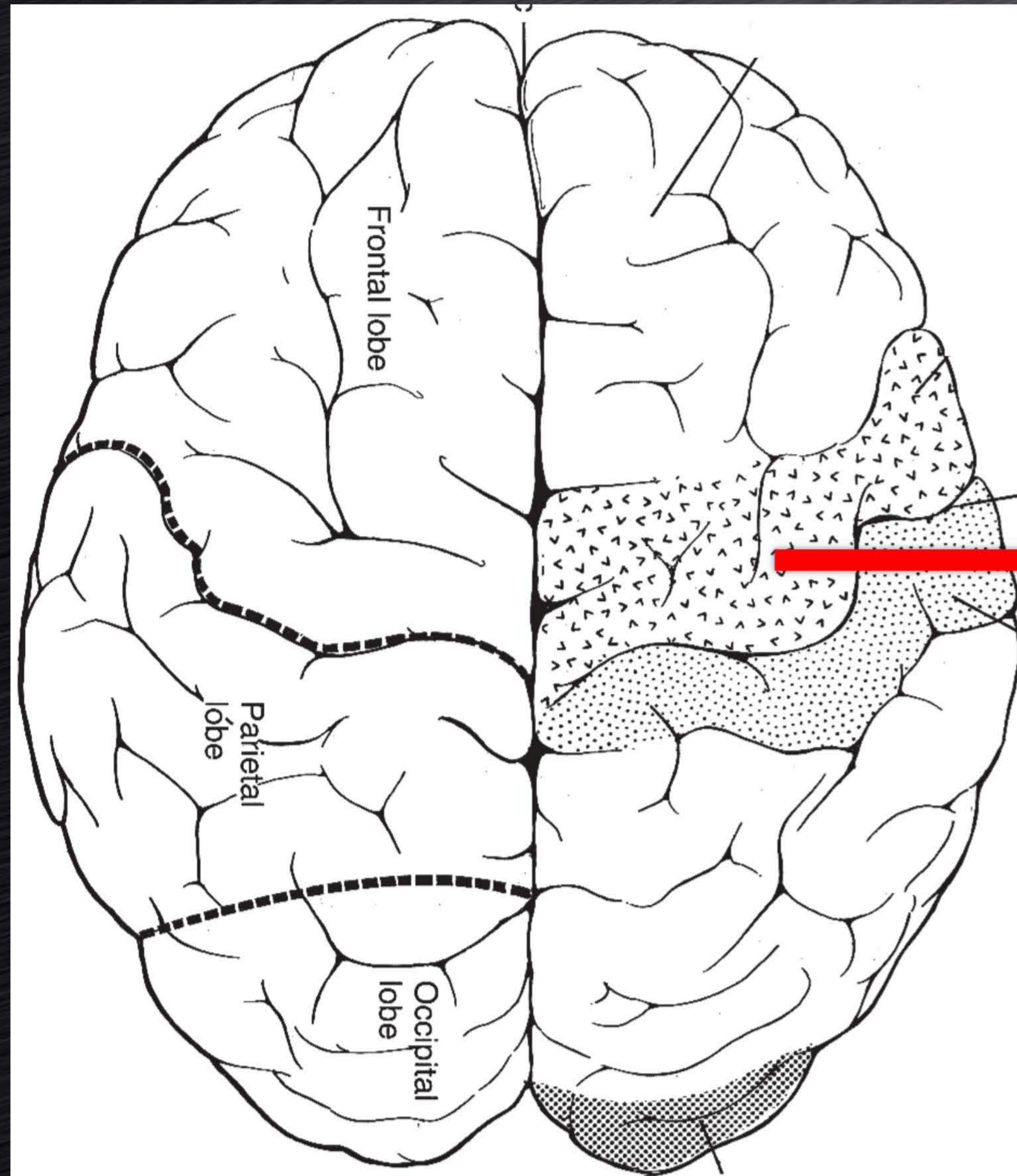
No problem.

I couldn't run my lab without BCI. I do molecular neuroscience research and my grant pays three people.

I'm writing this with my EEG courtesy of the Wadsworth Center Brain-Computer Interface Research Program ([www.wadsworth.org](http://www.wadsworth.org))

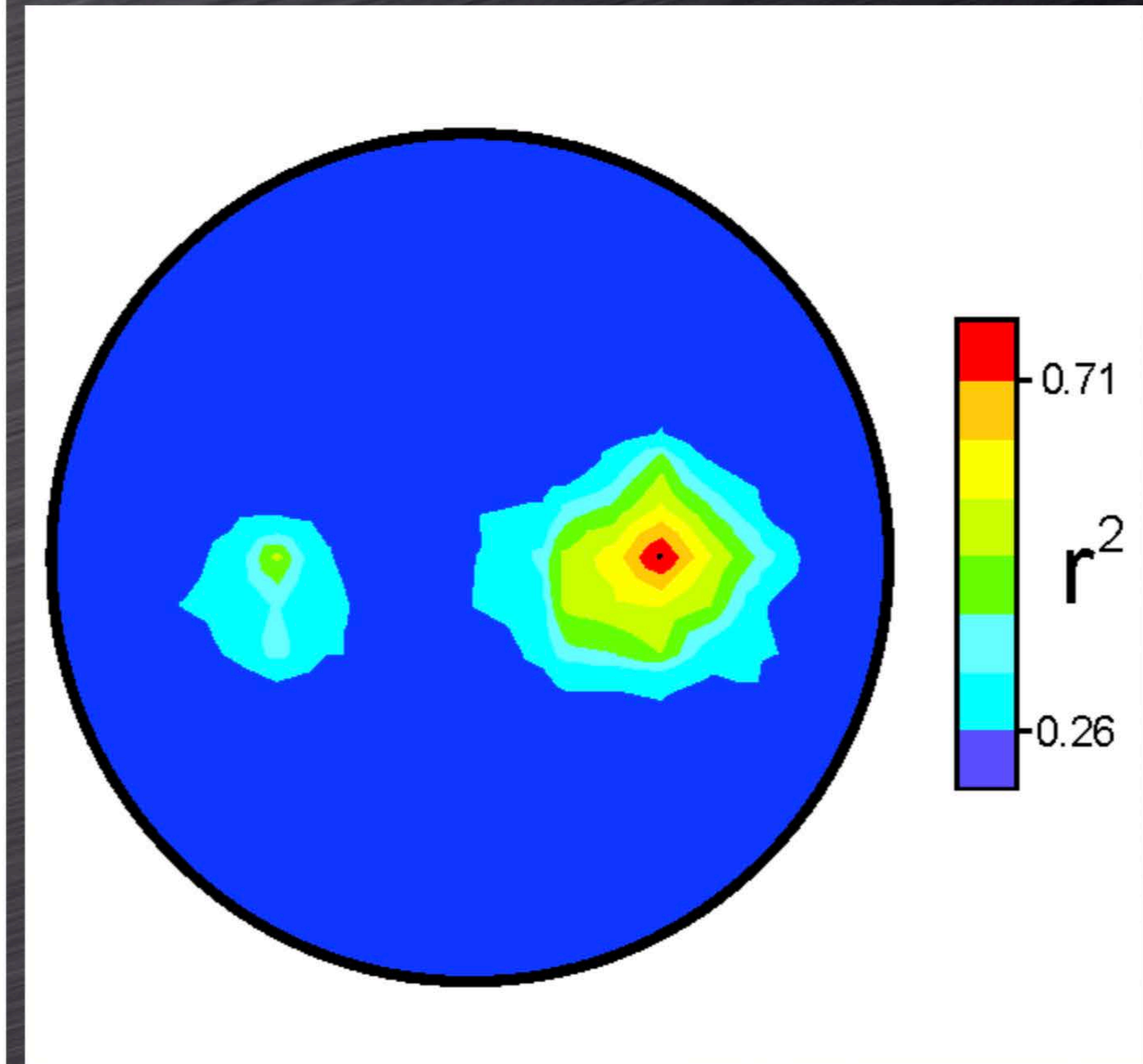
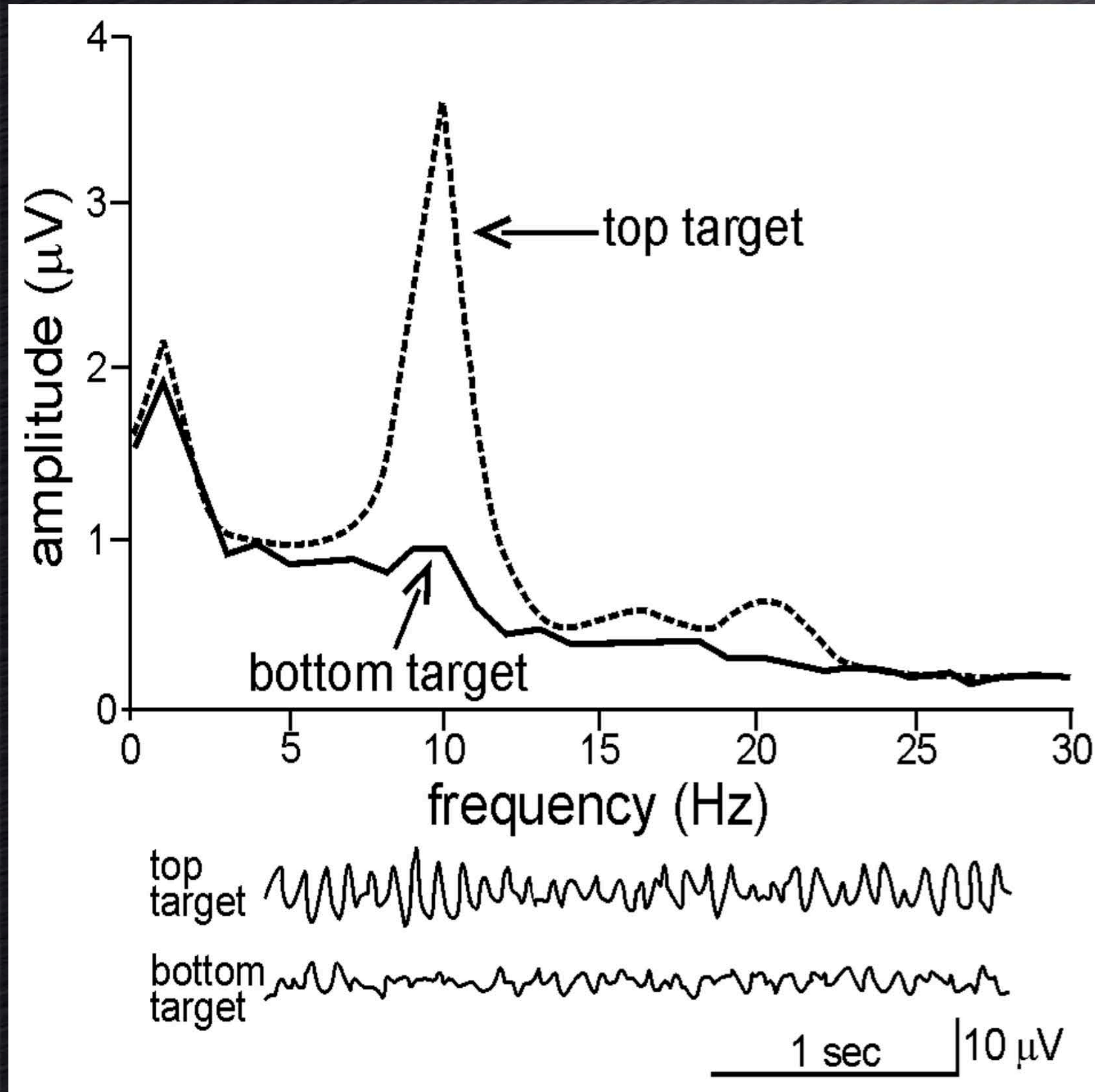


# EEG Sensorimotor Rhythms





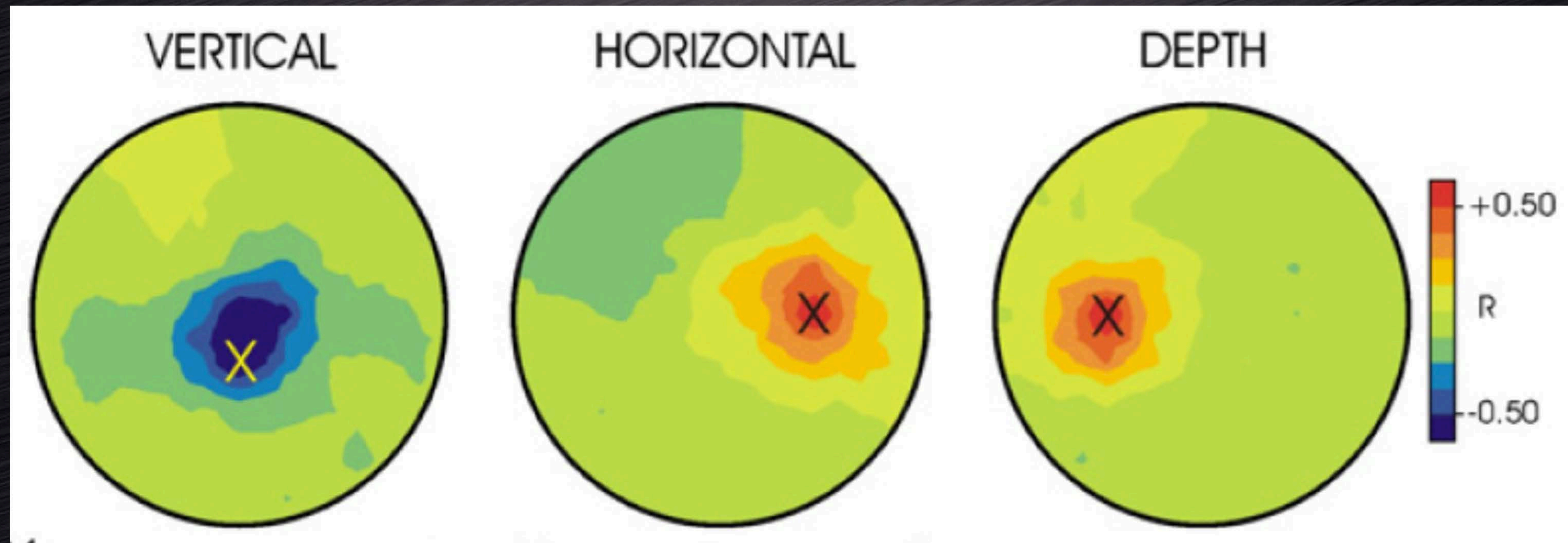
# EEG Sensorimotor Rhythm (SMR) Control: Spectral & Topographical Focus



Wolpaw/McFarland et al, Pfurtscheller et al, Kostov/Pollack, Penny et al



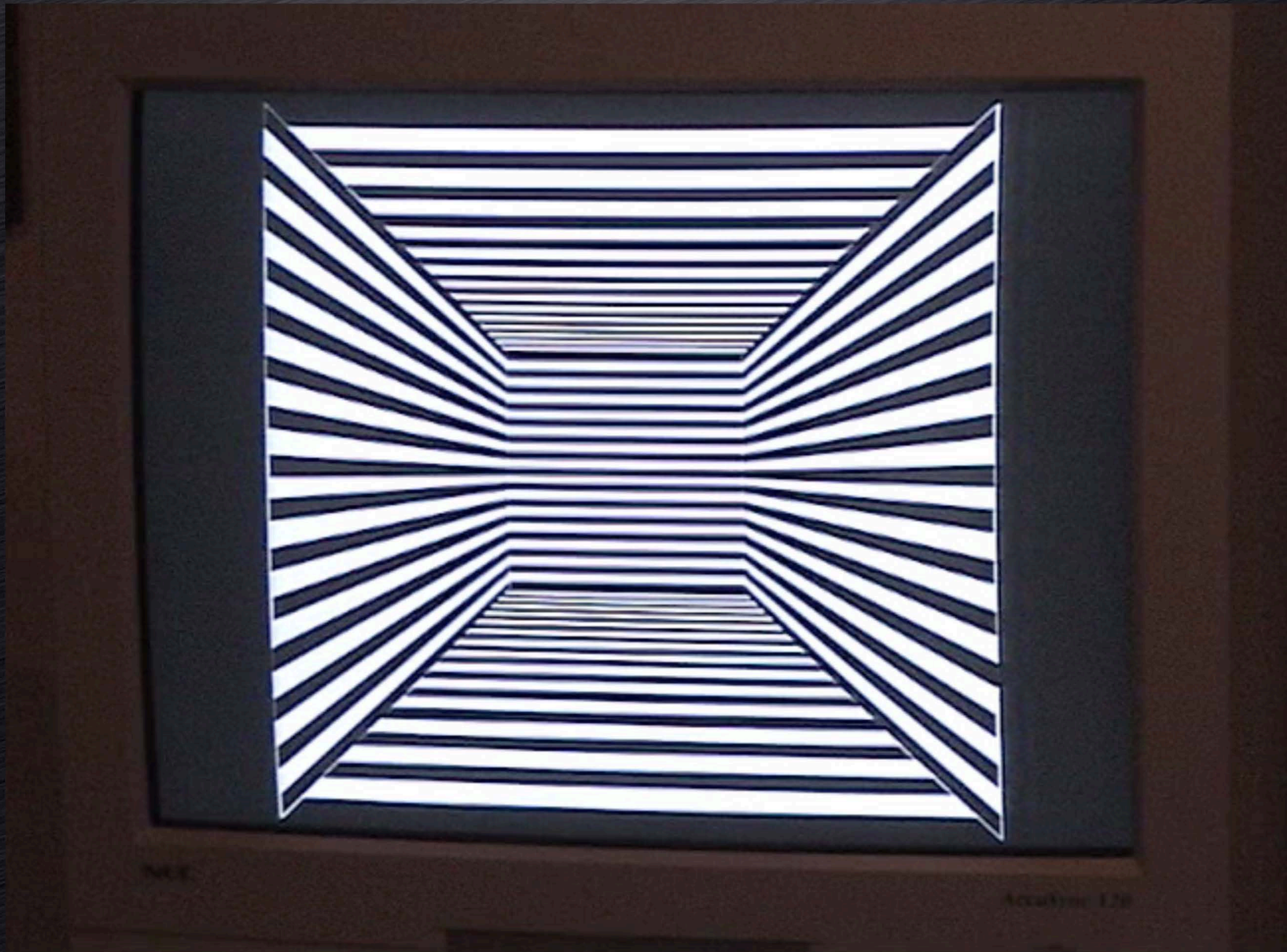
# Simultaneous Independent Channels of SMR Control



McFarland et al. JNE 2010



# Three-Dimensional Movement Control with SMRs





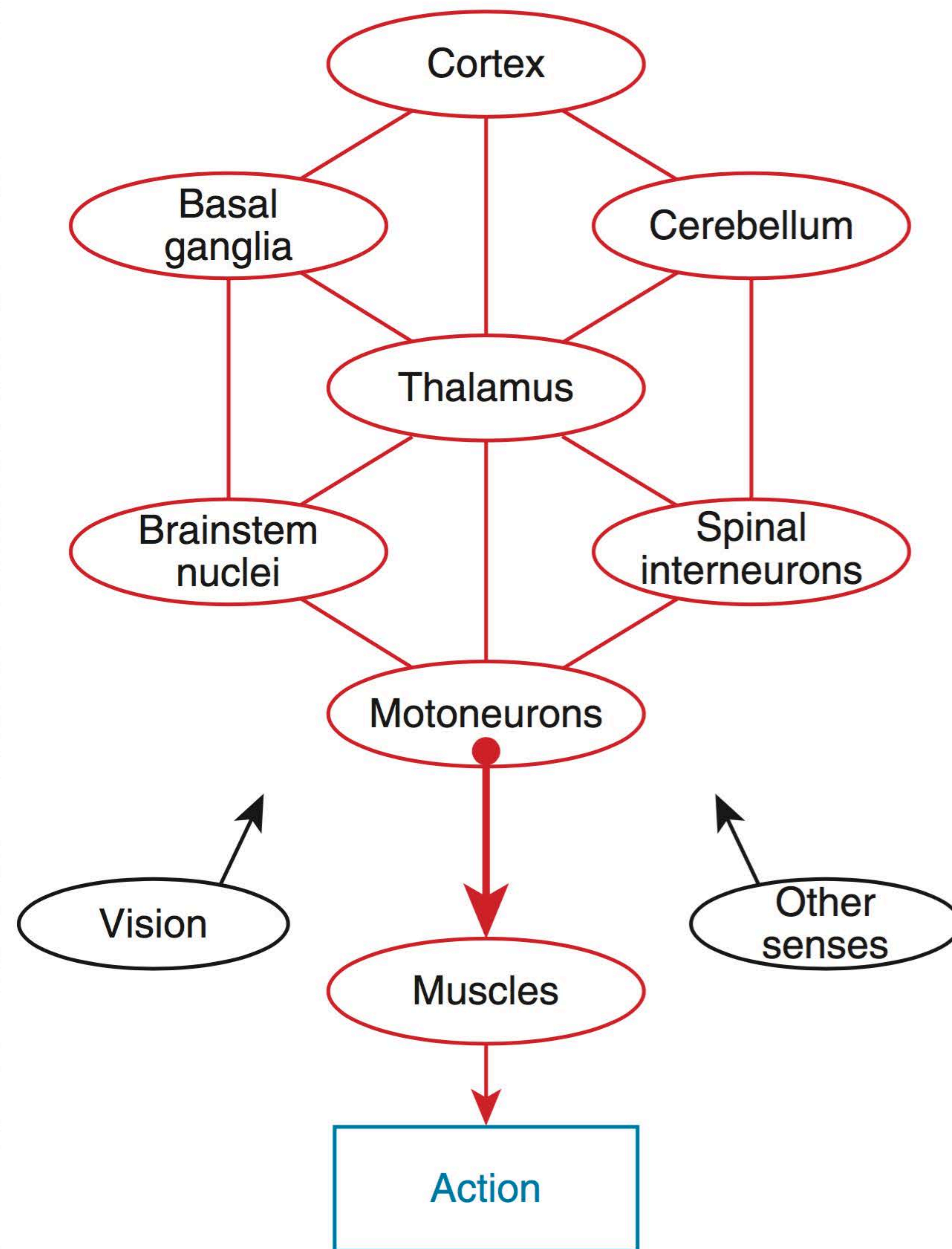
# The Grand Canyon Problem



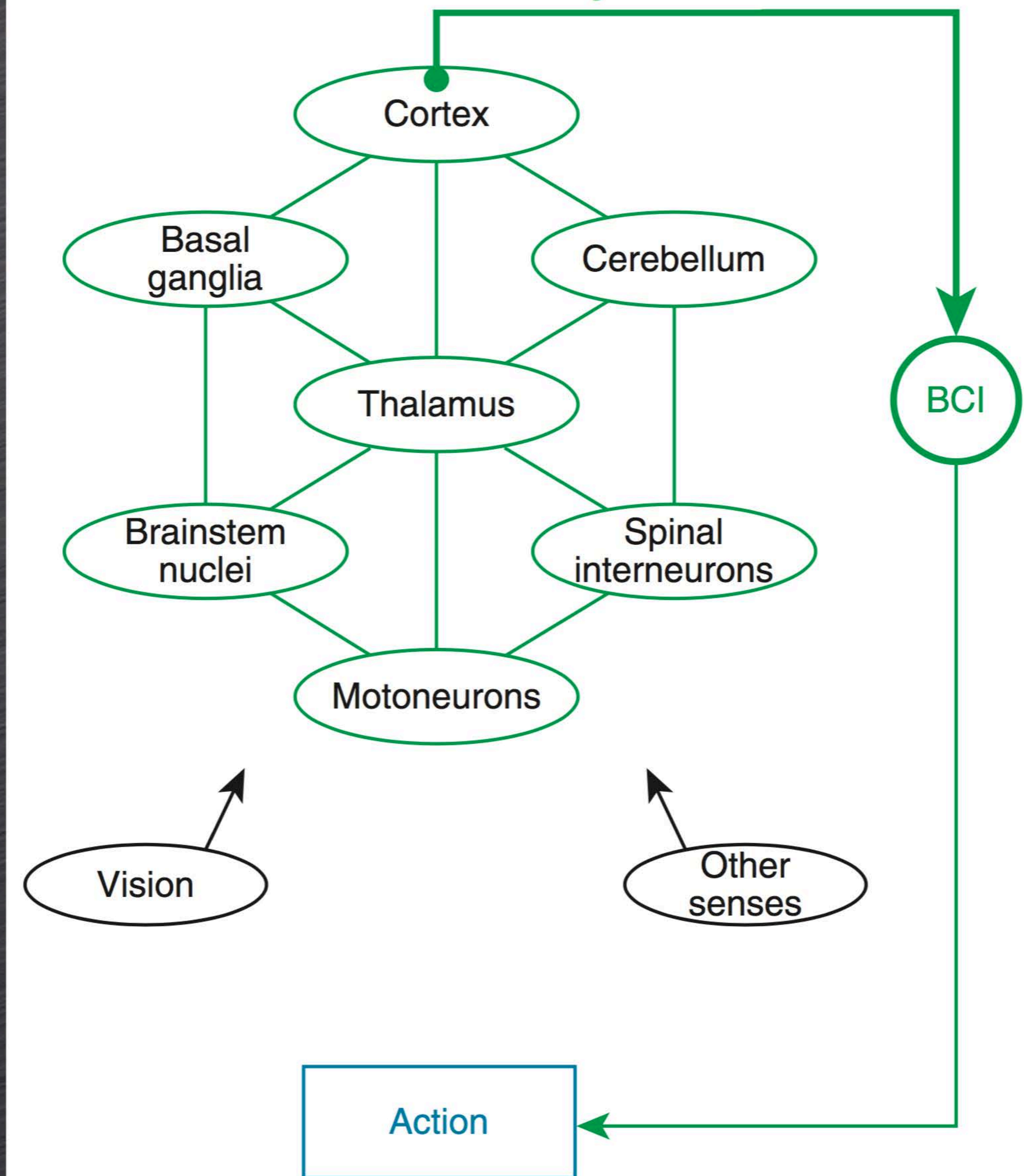


# Primary Origin of the Grand Canyon Problem

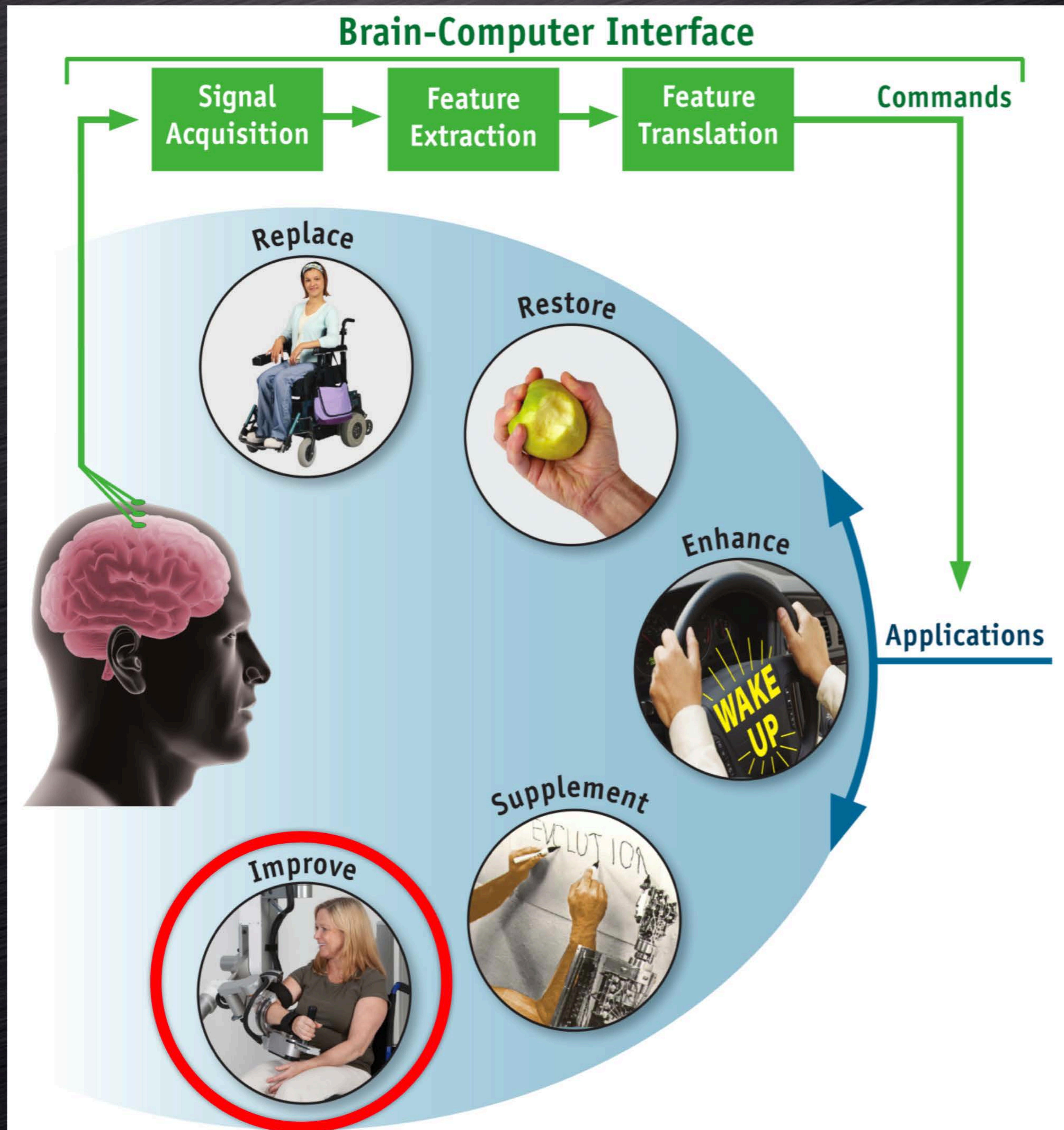
## CNS/Muscle System



## CNS/BCI System



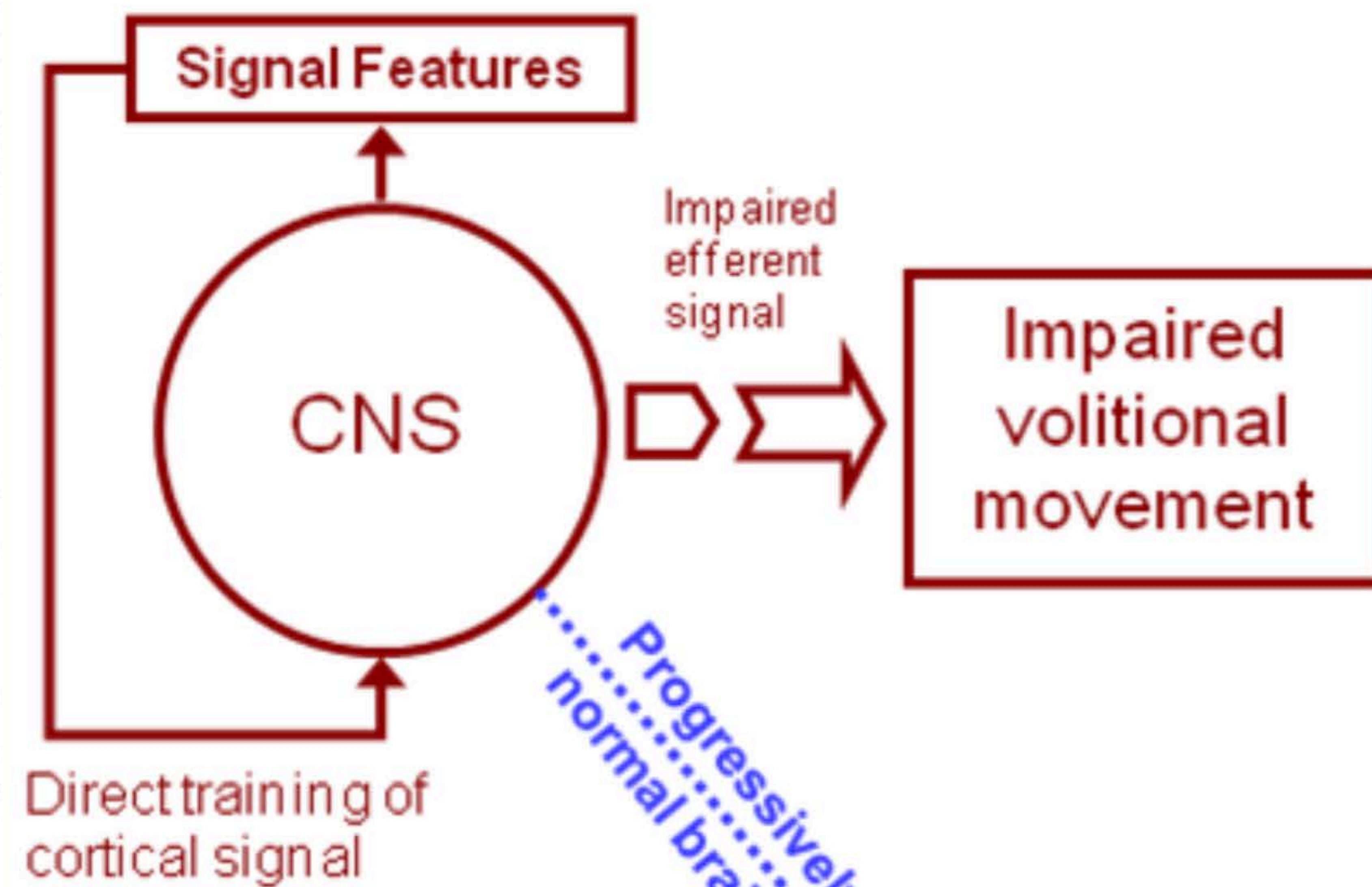




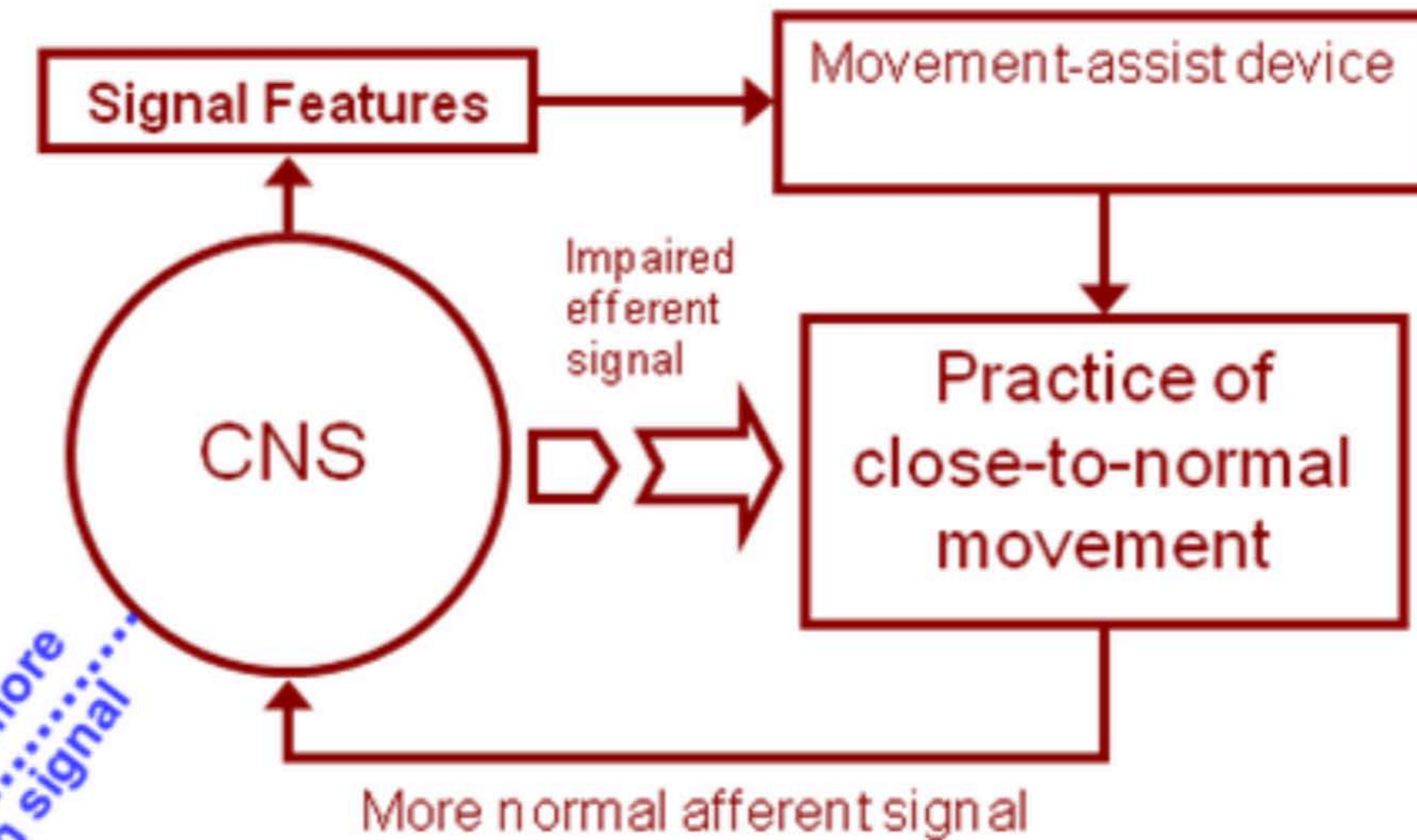


# BCI-based Rehabilitation

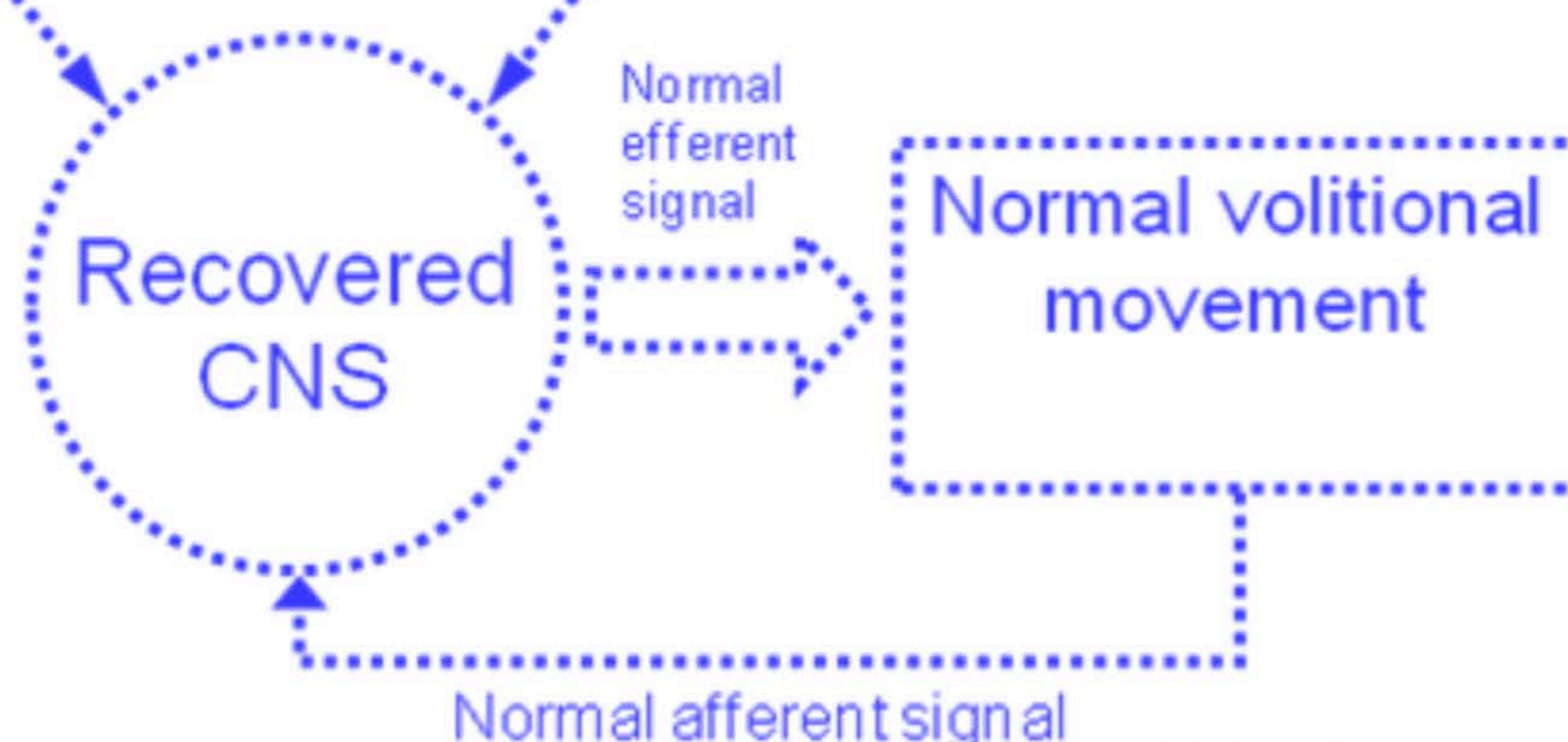
A. Training Strategy 1



B. Training Strategy 2

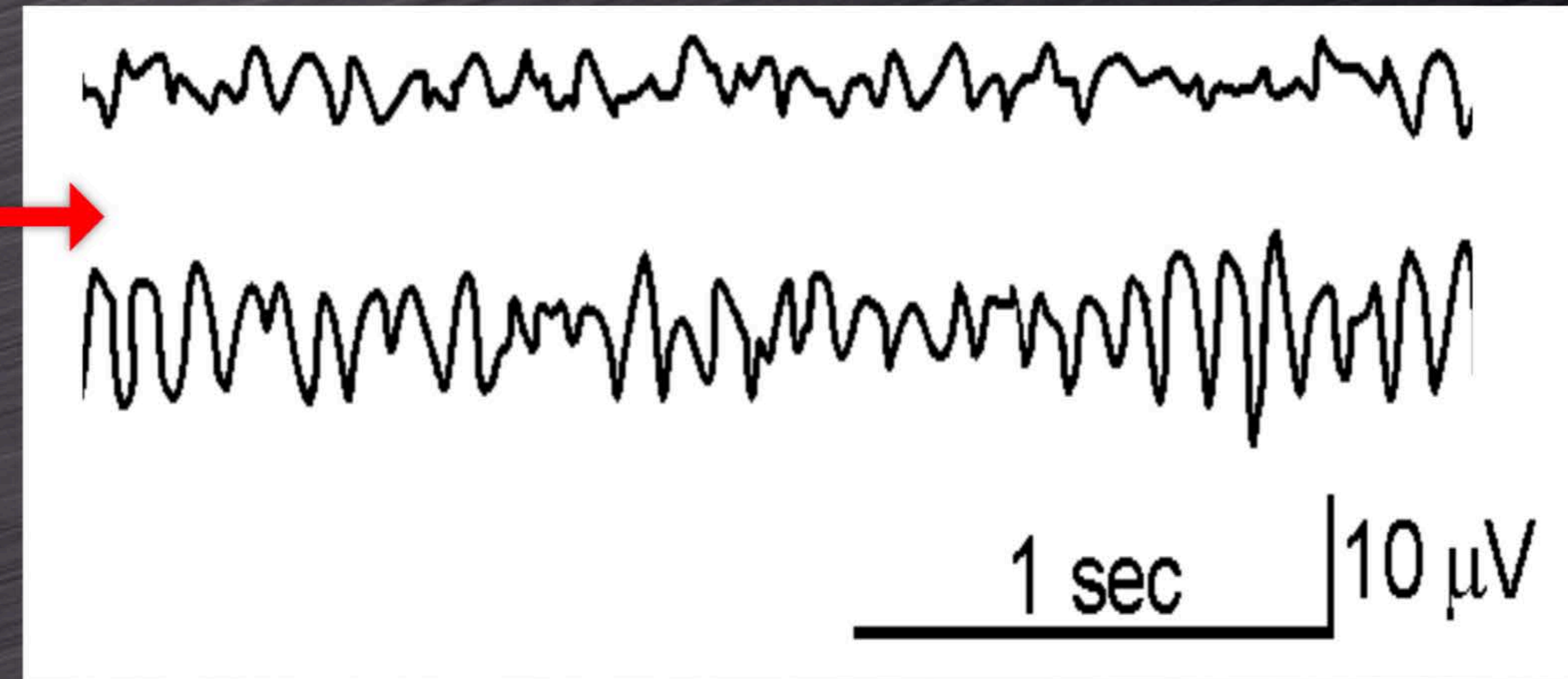
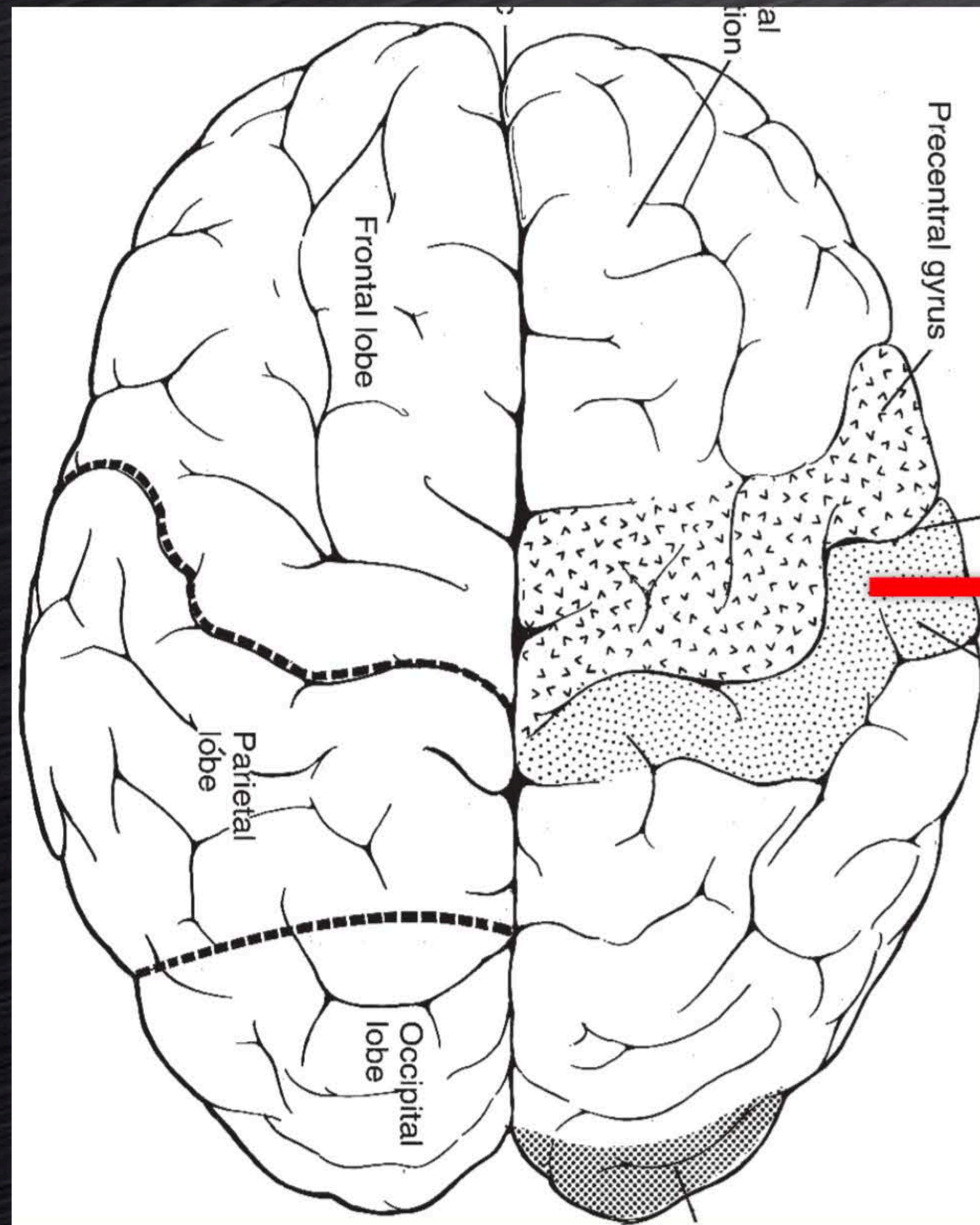


C. GOAL





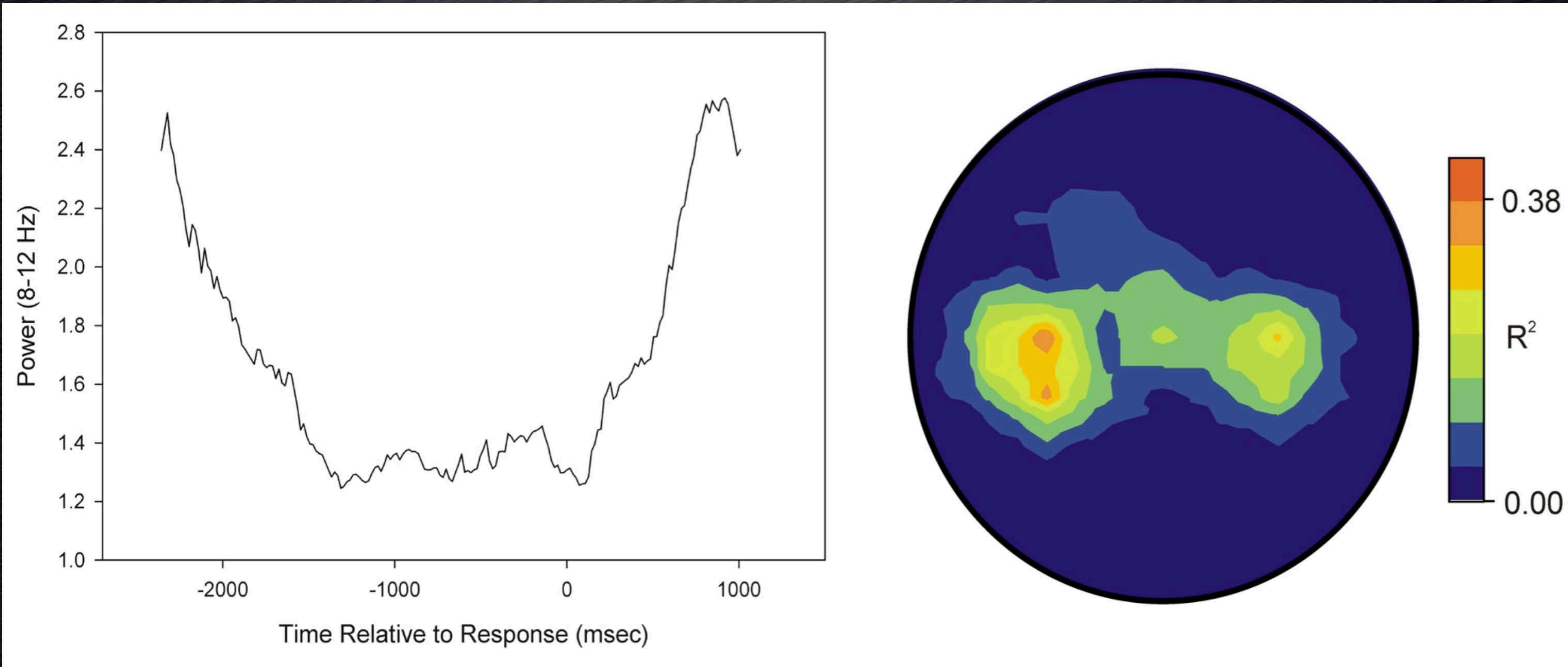
# *SMRs reflect the brain activity underlying movement*



McFarland et al. 2013



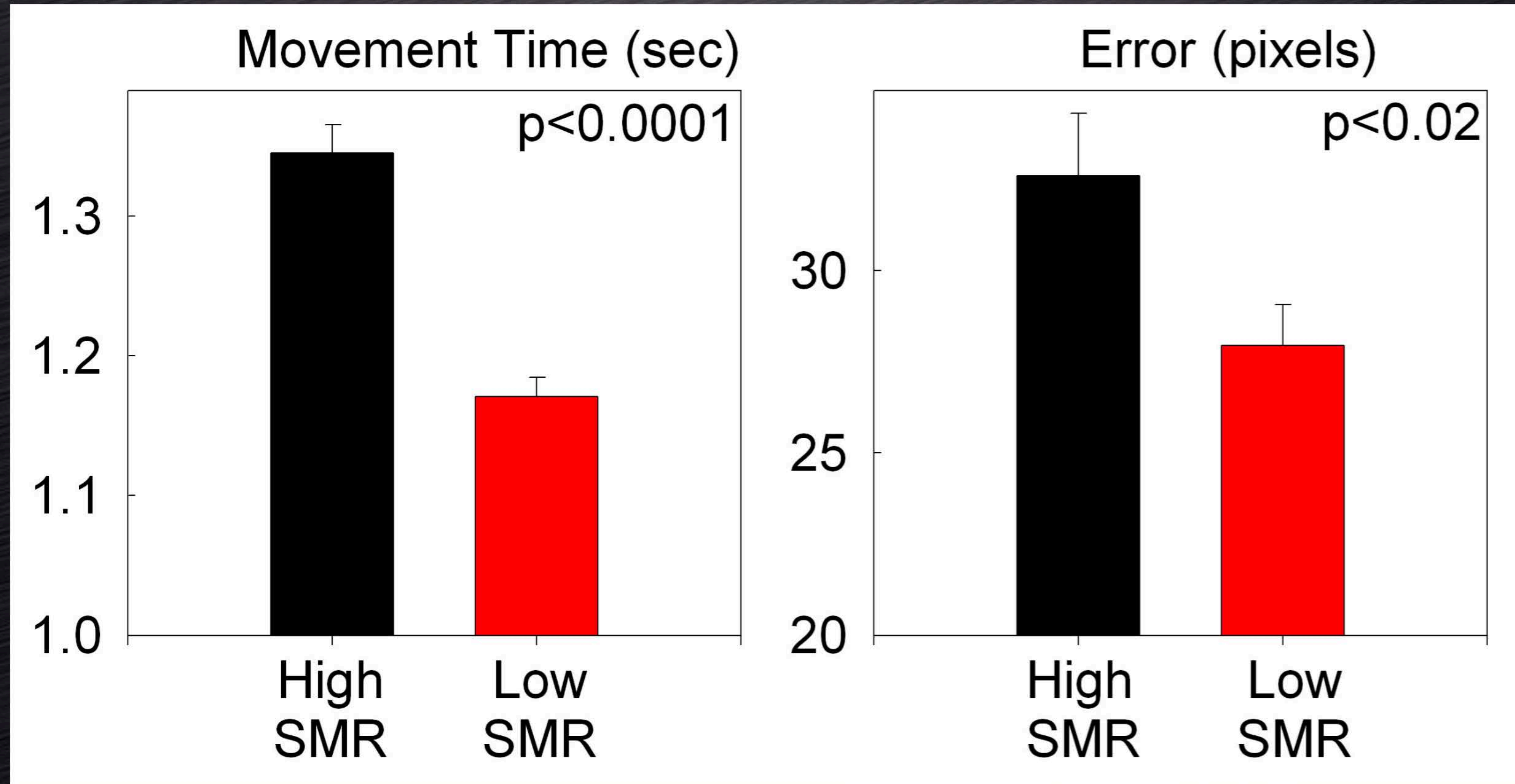
# *SMRs reflect the brain activity underlying movement*



McFarland et al. 2013



# *SMRs reflect the brain activity underlying movement*

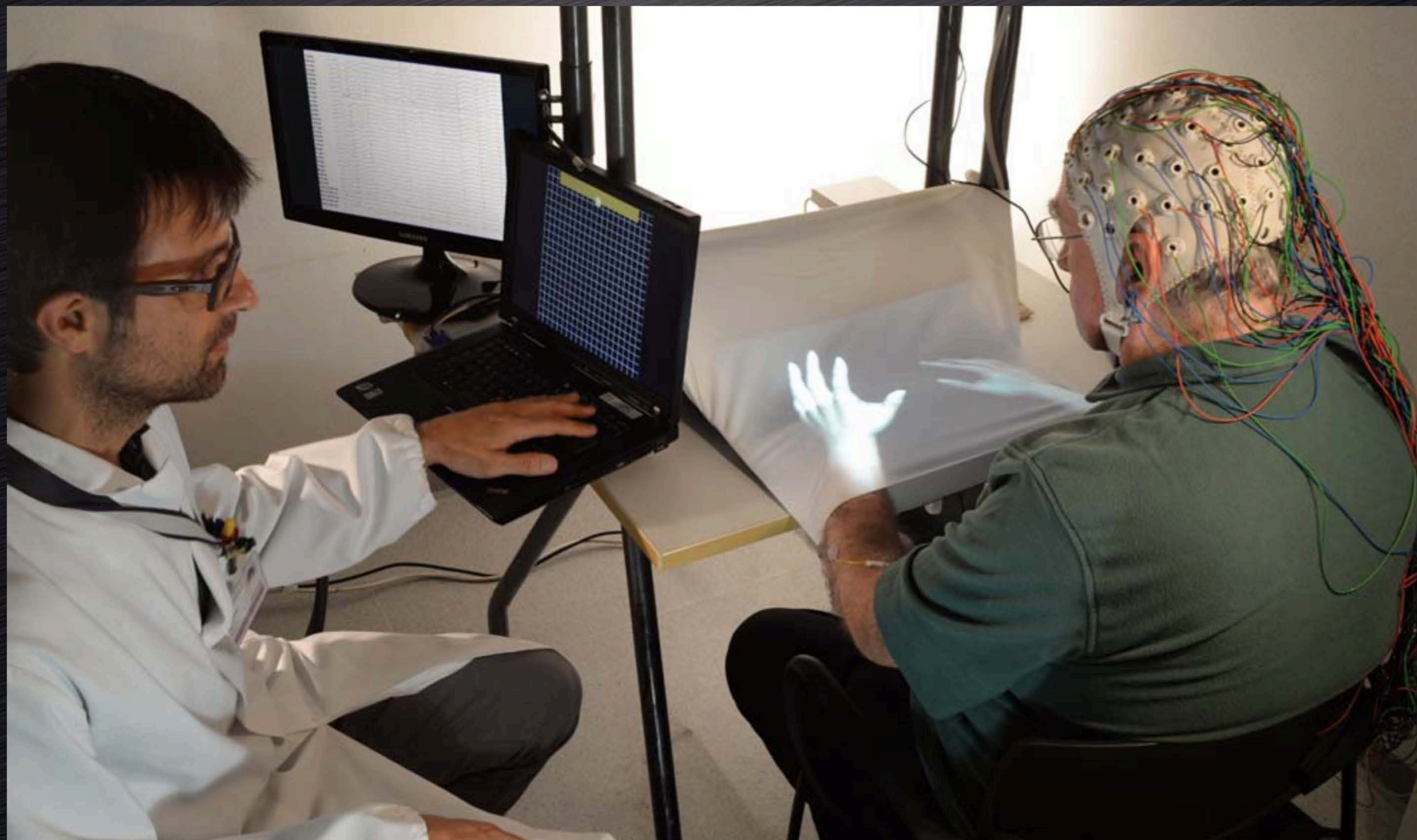


*Modifying them might improve impaired movement*

McFarland et al. 2013



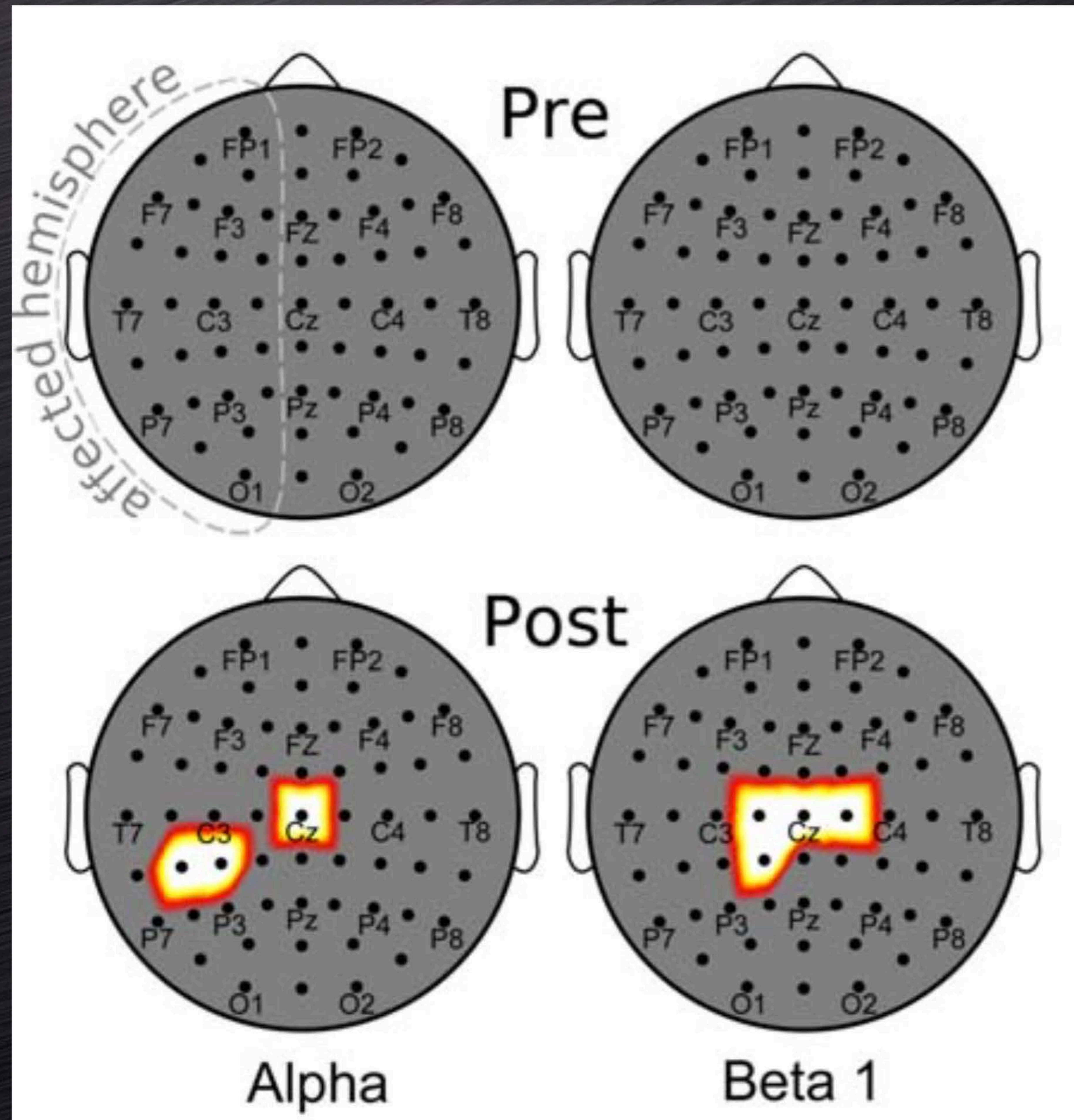
# Brain-Computer Interface Boosts Motor Imagery Practice during Stroke Recovery



Pichiorri et al. Ann Neurol 2015



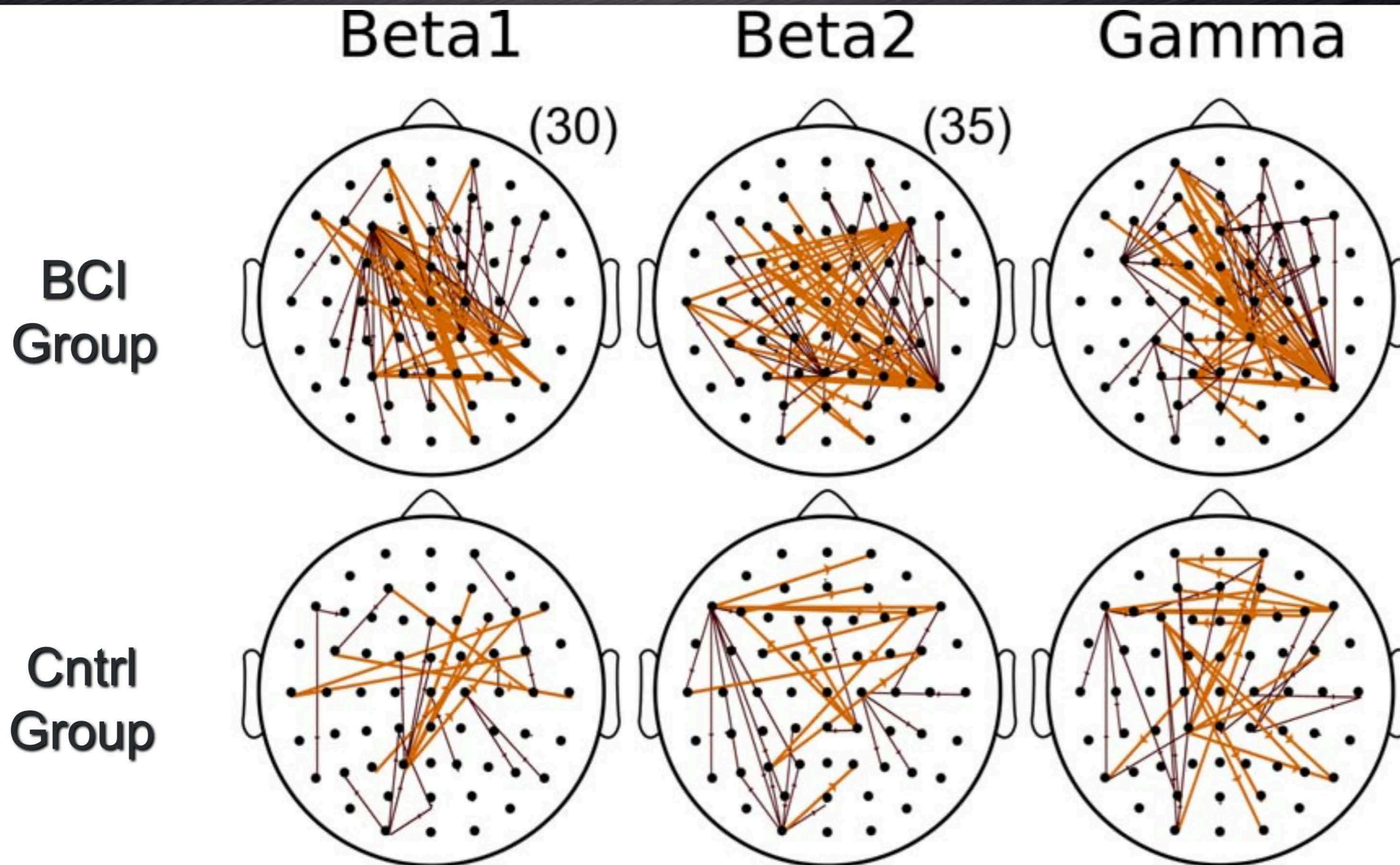
# BCI-based Stroke Rehabilitation



Pichiorri et al. Ann Neurol 2015

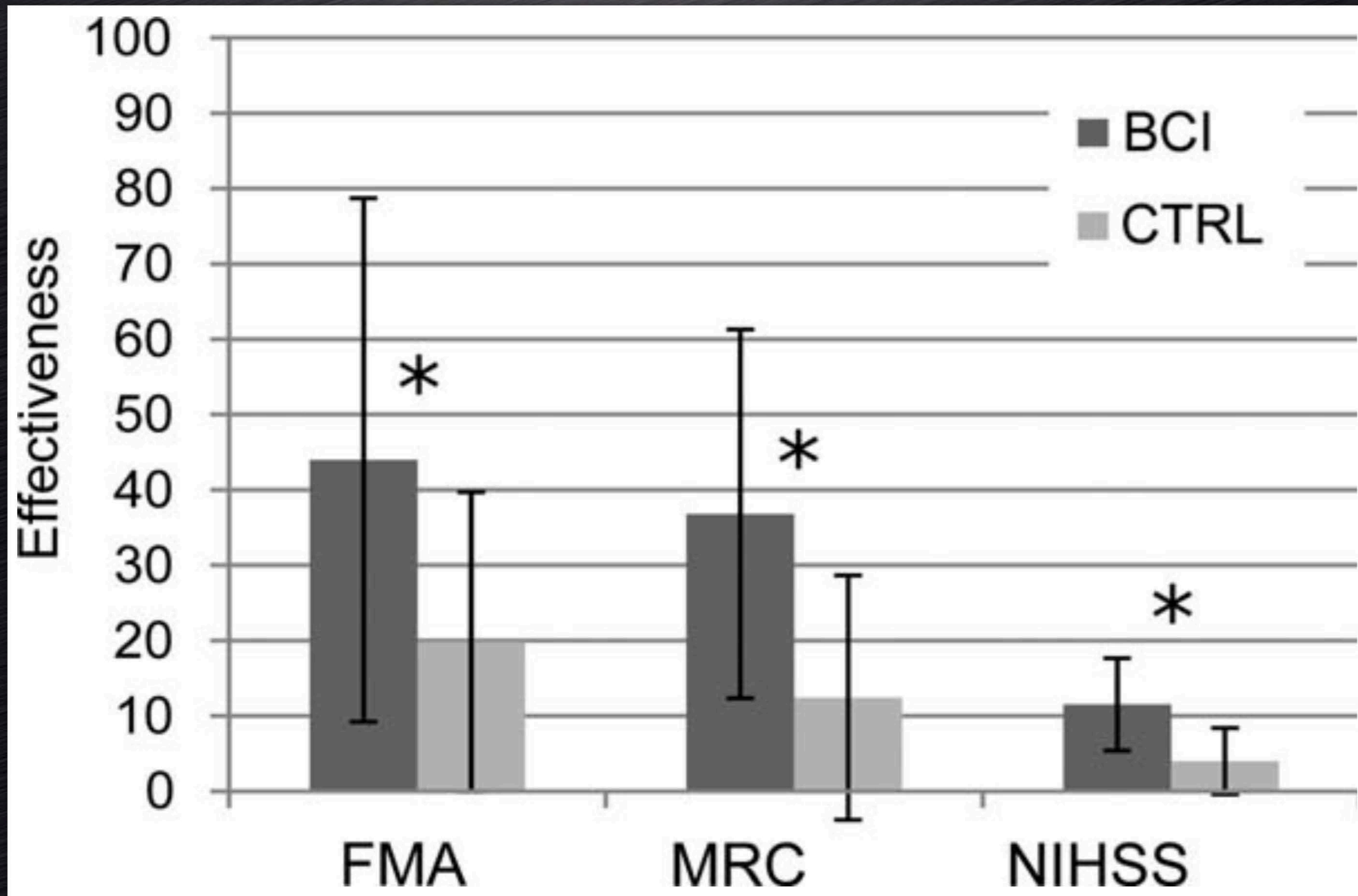


# BCI-based Stroke Rehabilitation





# BCI-based Stroke Rehabilitation



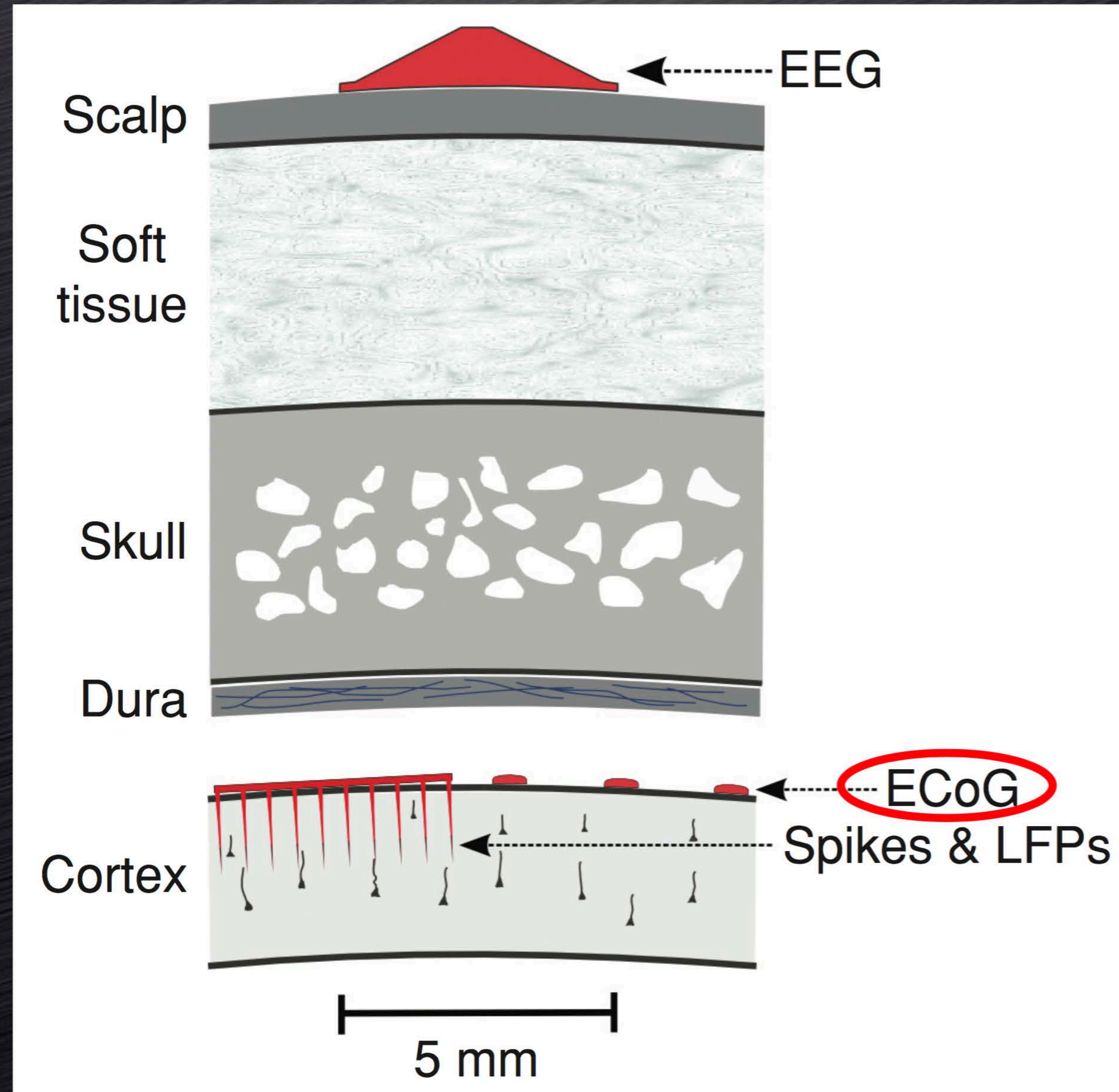
Pichiorri et al. Ann Neurol 2015



*Using electrocorticography  
(ECoG) to localize cortical  
function before surgery*



# Electrophysiological Imaging of Brain Activity





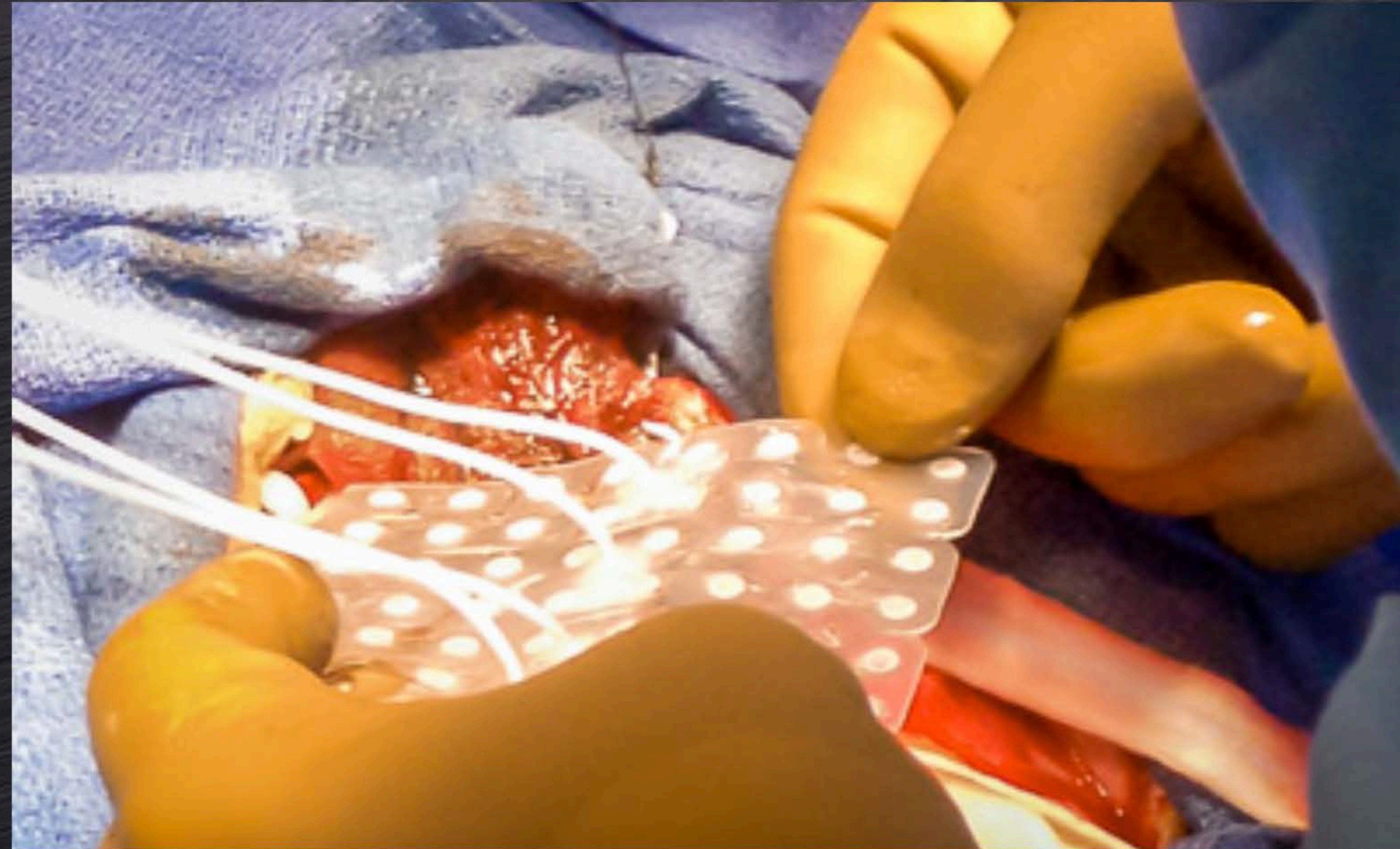
# Imaging Brain Processes: Requirements

	Spatial Resolution	Temporal Resolution	Coverage
EEG	Red	Green	Green
fMRI	Green	Red	Green
Intracortical	Green	Green	Red
ECoG	Green	Green	Green



# ECoG-based Functional Mapping prior to Surgery

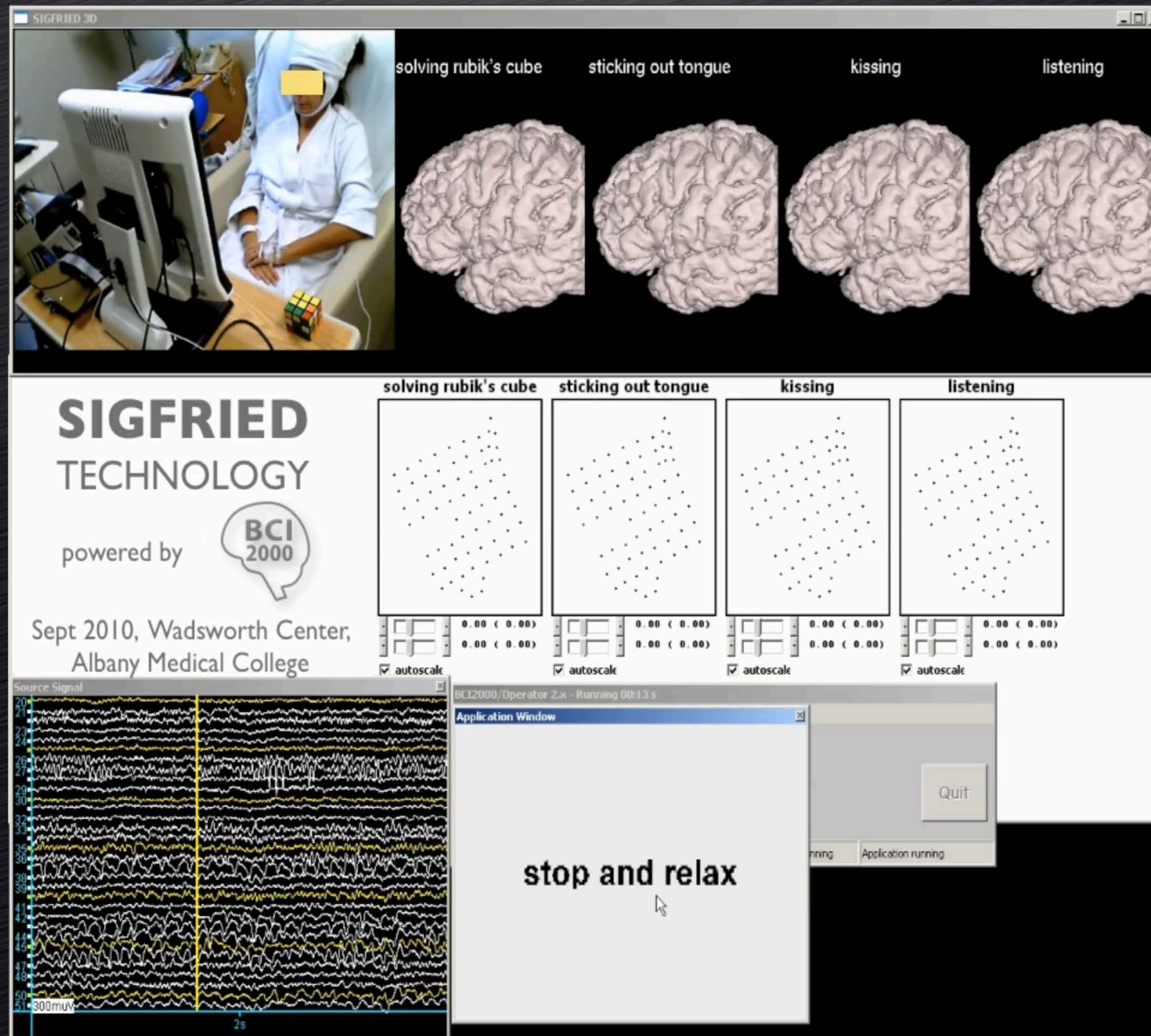
- Epilepsy surgery often requires functional mapping



- Traditional stimulation method is time-consuming and risky
- ECoG-based passive mapping is a promising alternative
- New mapping method (SIGFRIED; Schalk et al. 2008)

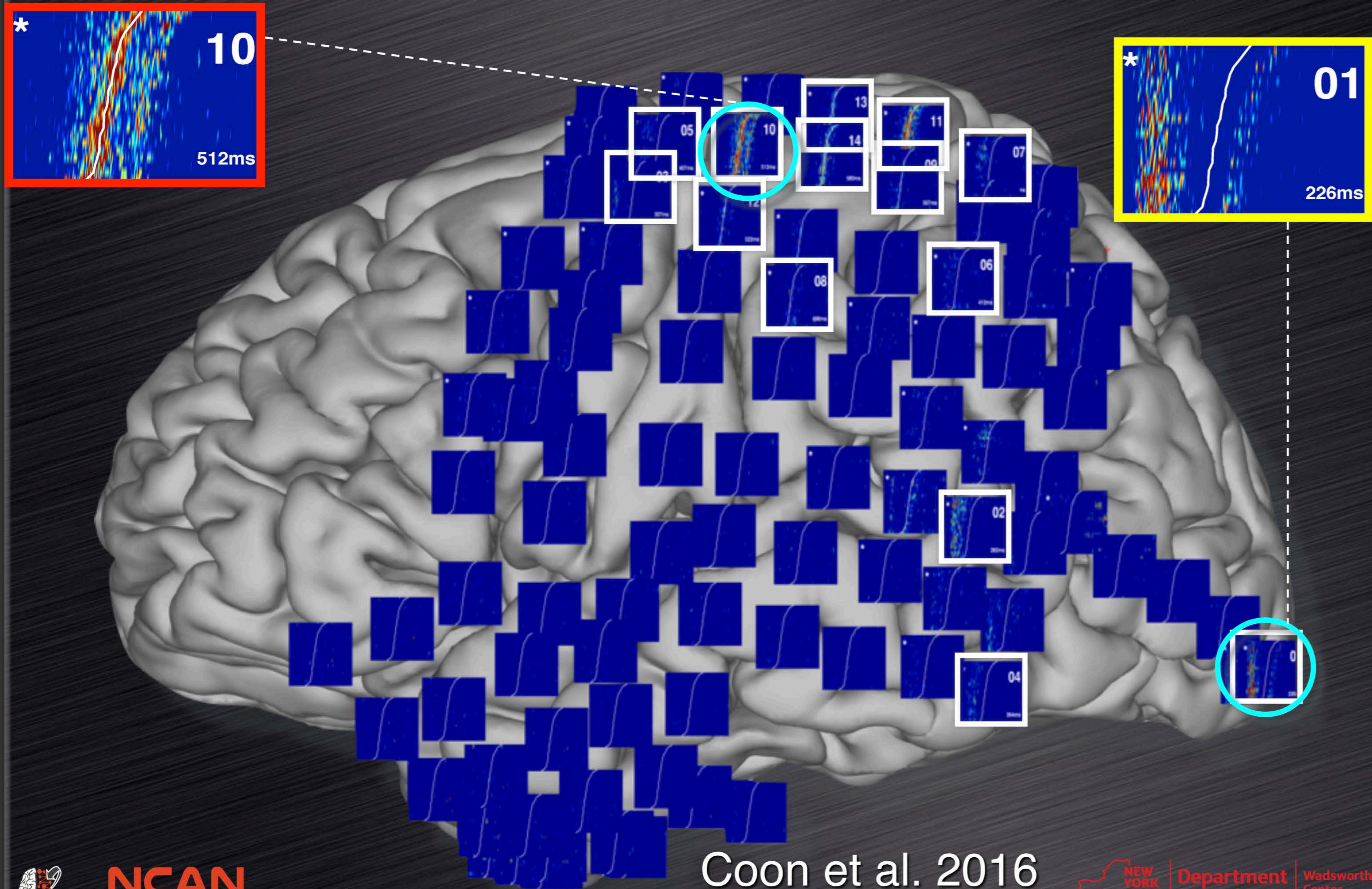


# Real-Time Functional Imaging





# The Sequential Activity Underlying a Behavior





*Reflex conditioning for rehabilitation  
after spinal cord injury:  
Targeted Neuroplasticity*

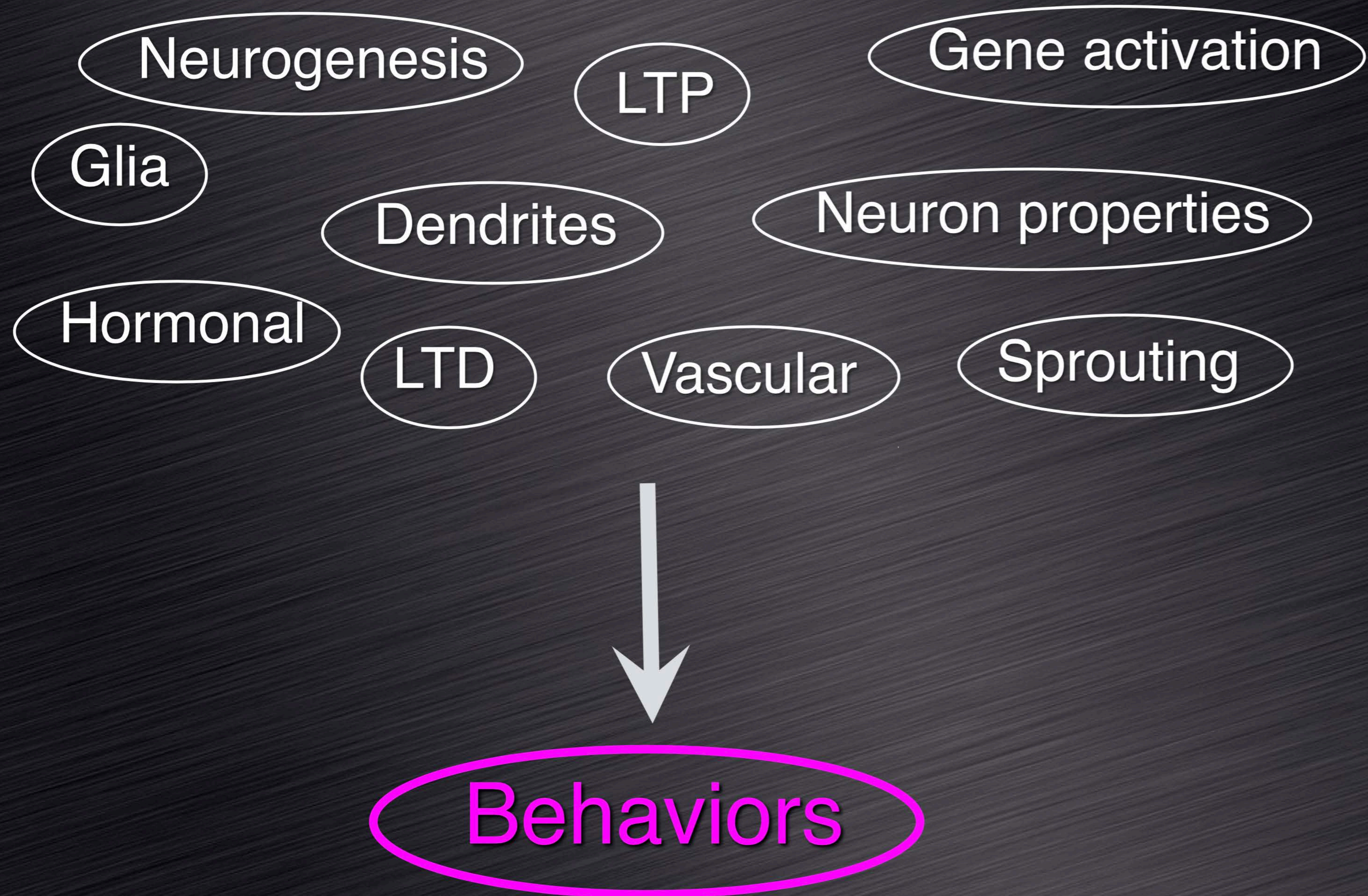


# Traditional Rehabilitation

- Starts from the assumption that the CNS is largely hard-wired with little capacity for plasticity
- Tries to maximize remaining capacities and to induce whatever minimal plasticity is possible
- Two major strategies
  - Practice impaired behaviors (e.g., locomotor training)
  - Drugs to reduce dysfunction (e.g., baclofen for spasticity)
- Often limited efficacy
  - Important behaviors are not fully restored
  - Significant side effects may occur



# Plasticity is Ubiquitous and Continuous in the CNS



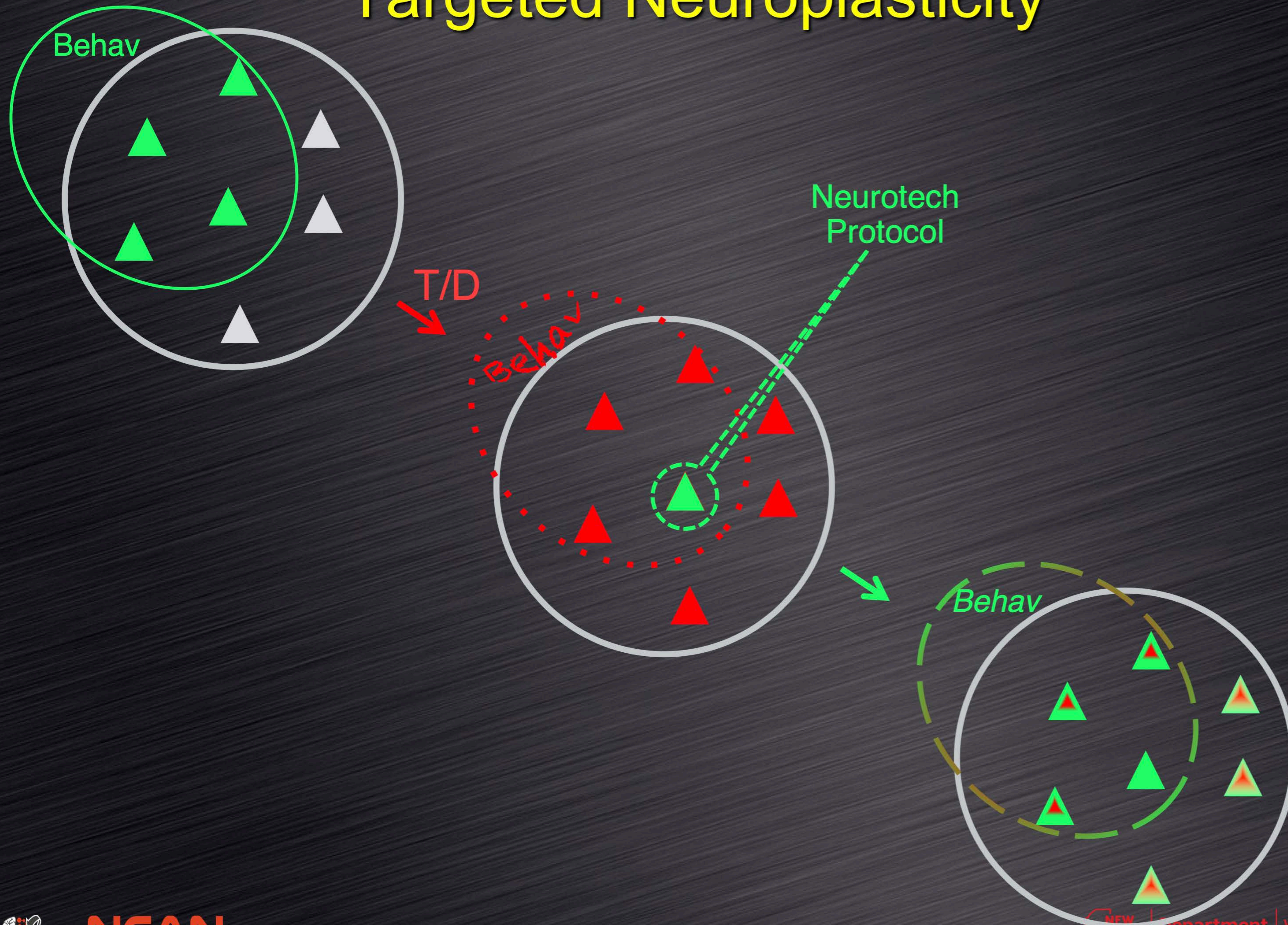


# Activity-Dependent Plasticity: Ubiquitous & Continuous in the CNS

- Many mechanisms;  $\gg 10^{12}$  sites; msec to years
- Thus, when function is impaired there are many options
- No direct overall guidance
- The drivers are local; most sites are far from behavior
- Thus, no certainty that optimal pattern of plasticity will occur
- Traditional rehab is not specific, does not target specific sites; seldom fully effective
- New methods are needed



# Targeted Neuroplasticity



**NCAN.**  
National Center for Adaptive Neurotechnologies

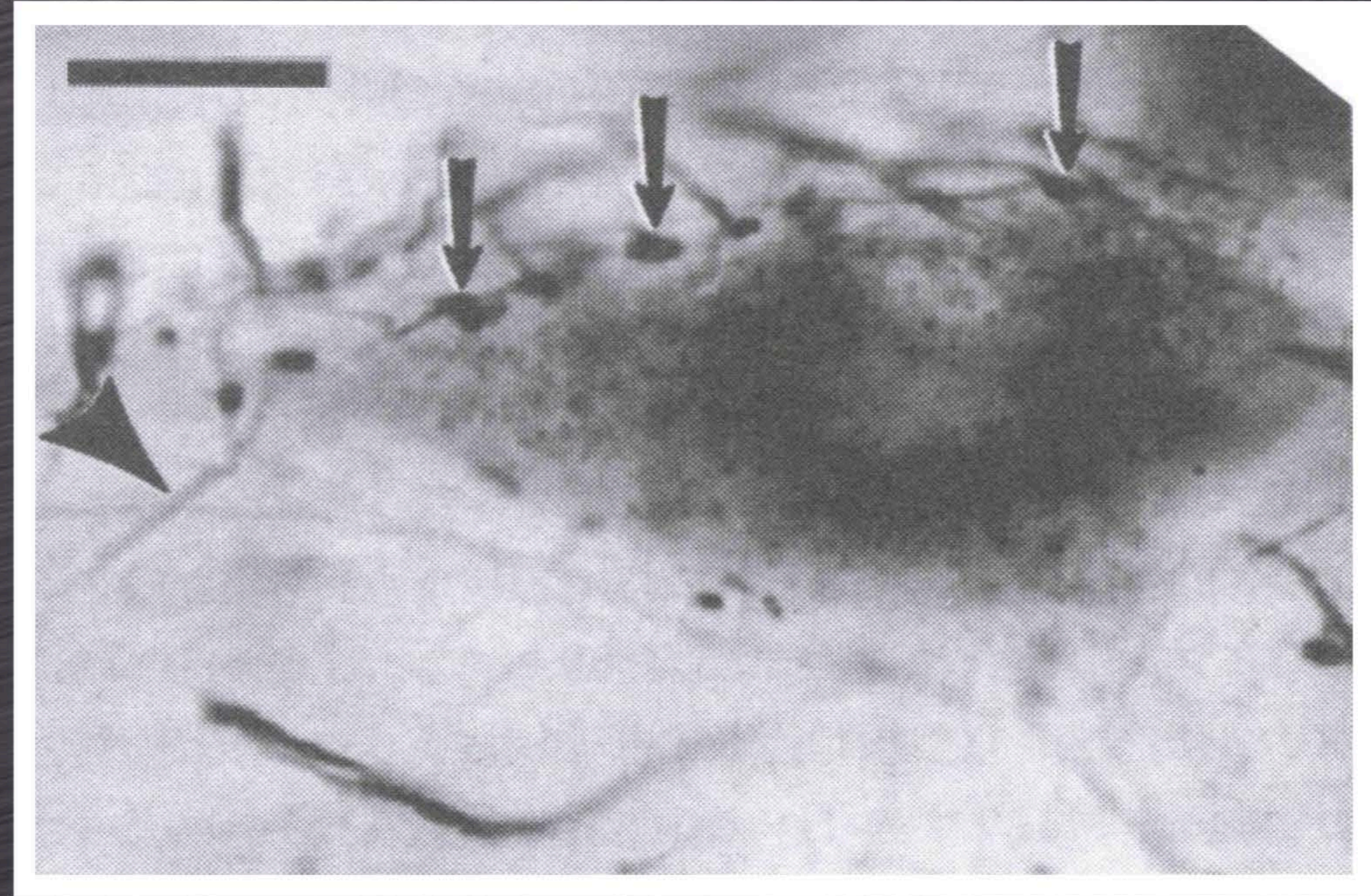
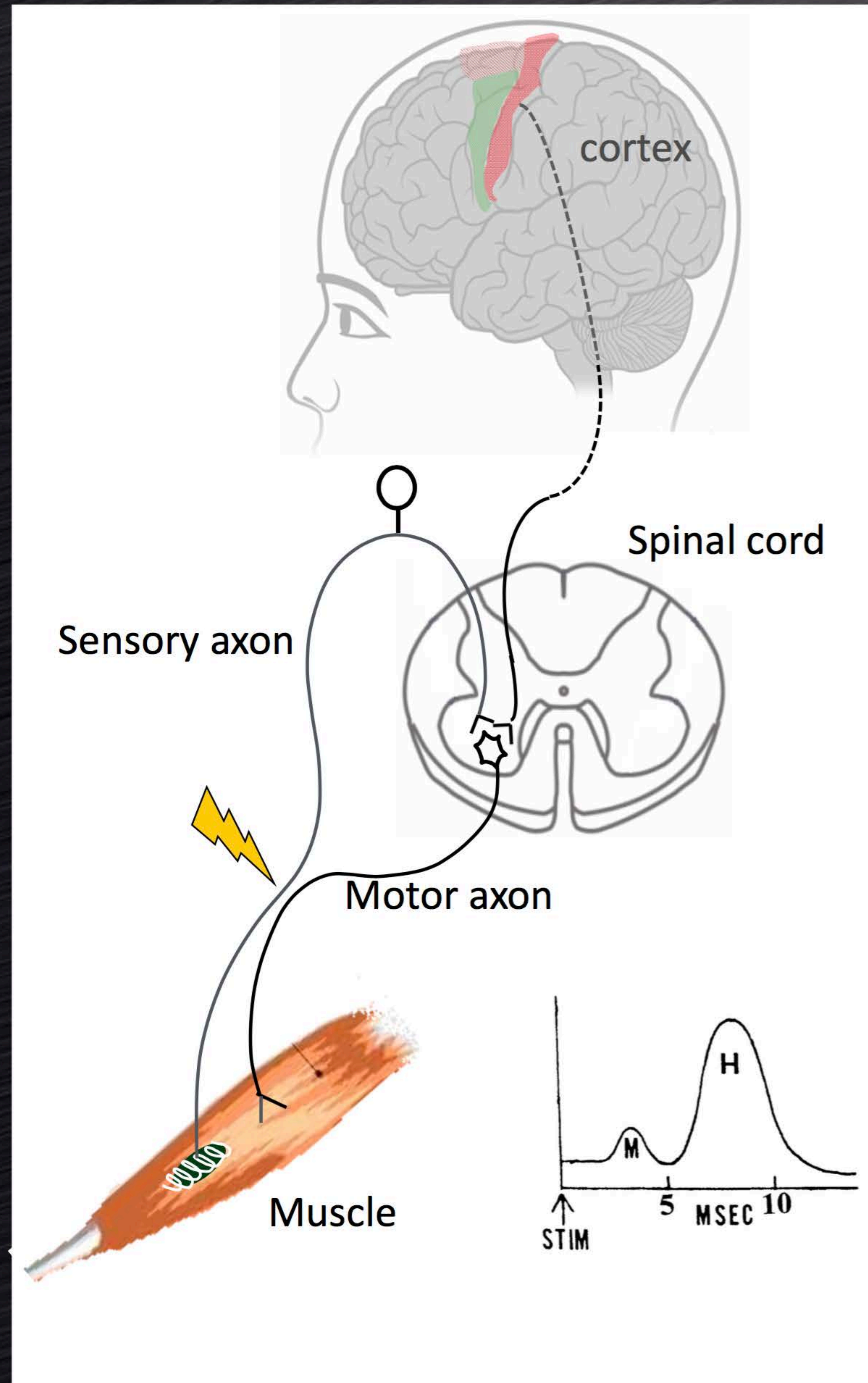


Department  
of Health

Wadsworth  
Center



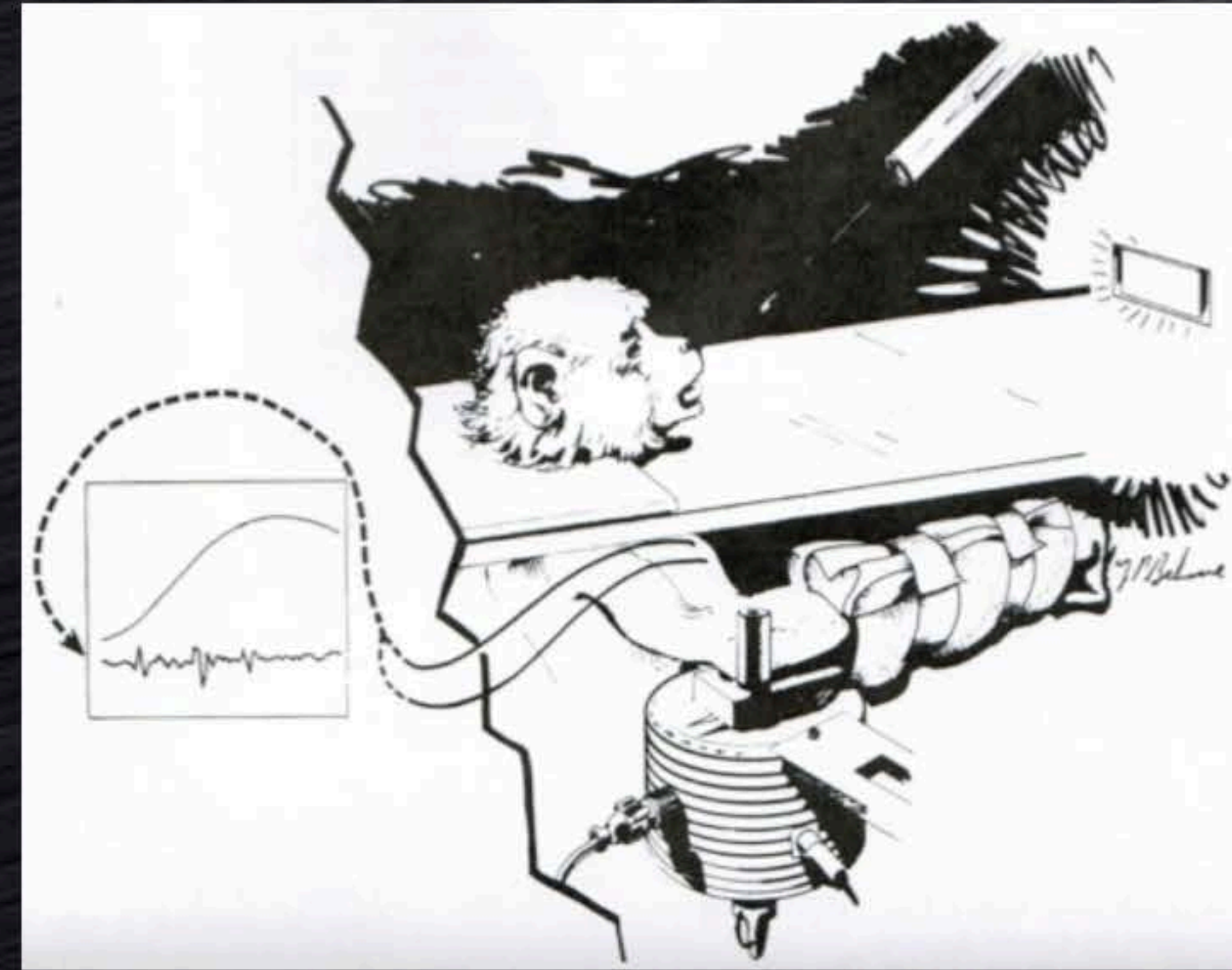
# Targeted Neuroplasticity: Modifying the H-Reflex



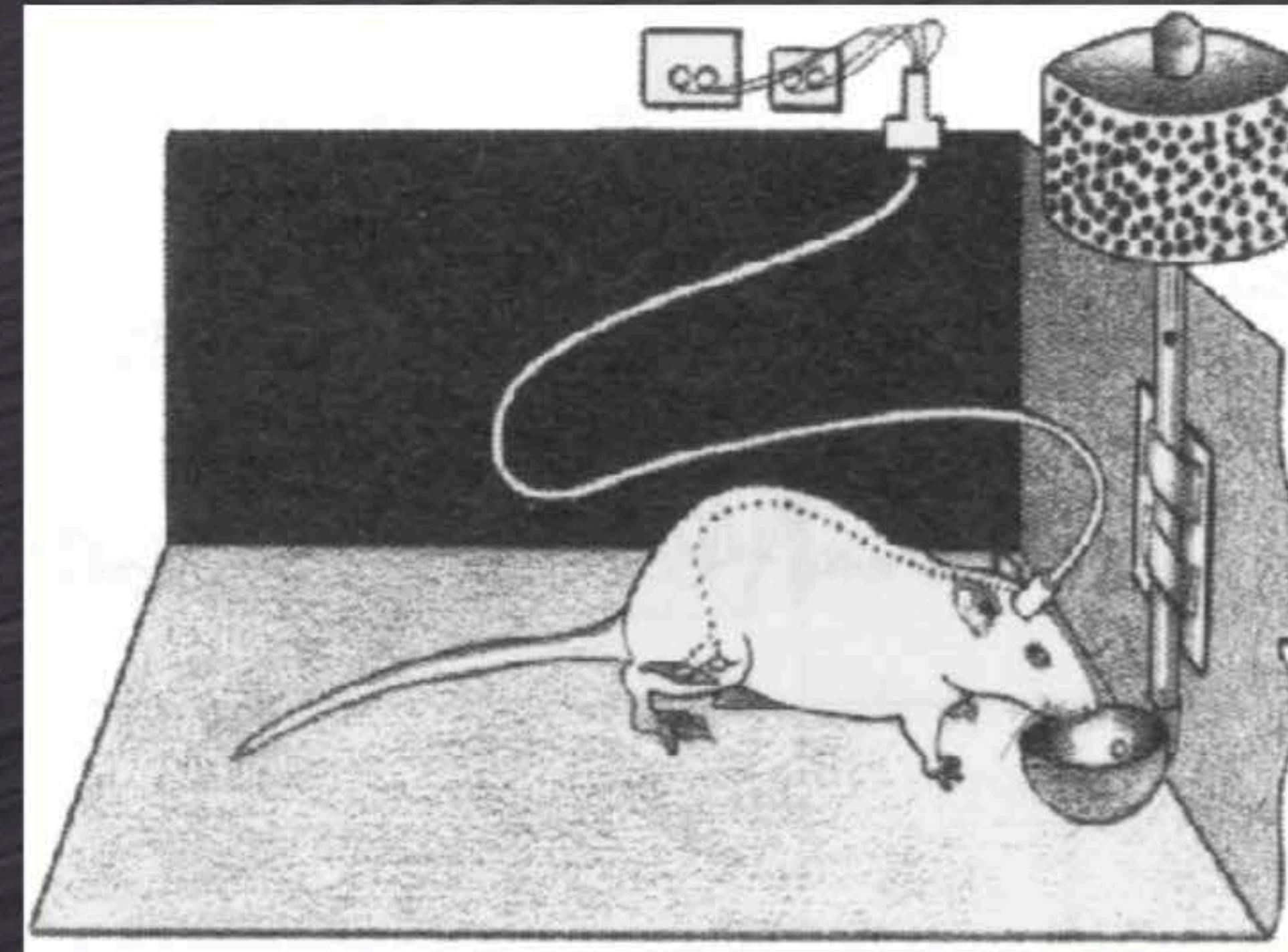
Monkeys, rats, mice, and people can gradually change H-reflex size when rewarded for doing so.



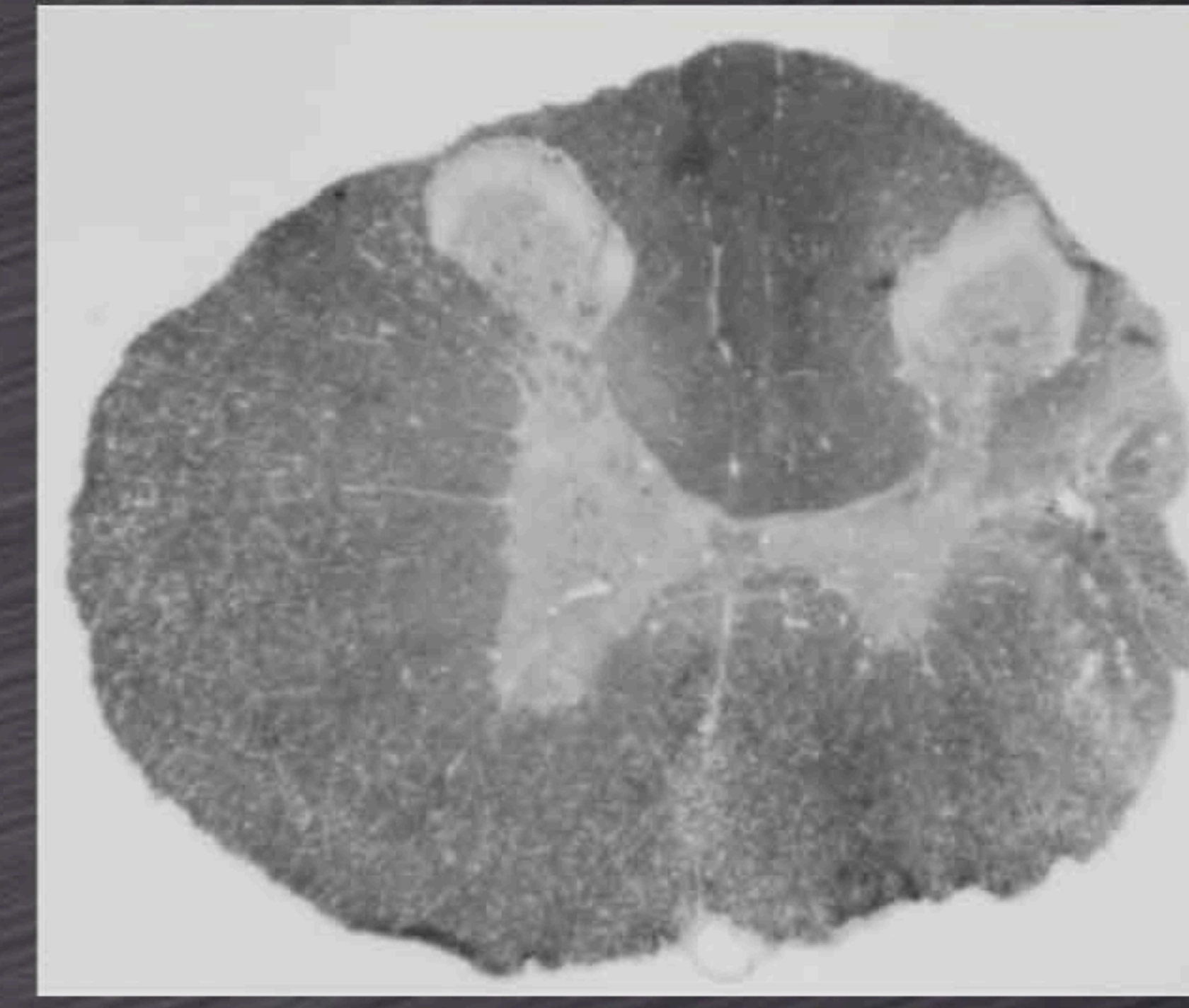
# Operant Conditioning of Spinal Reflexes 1978-2017



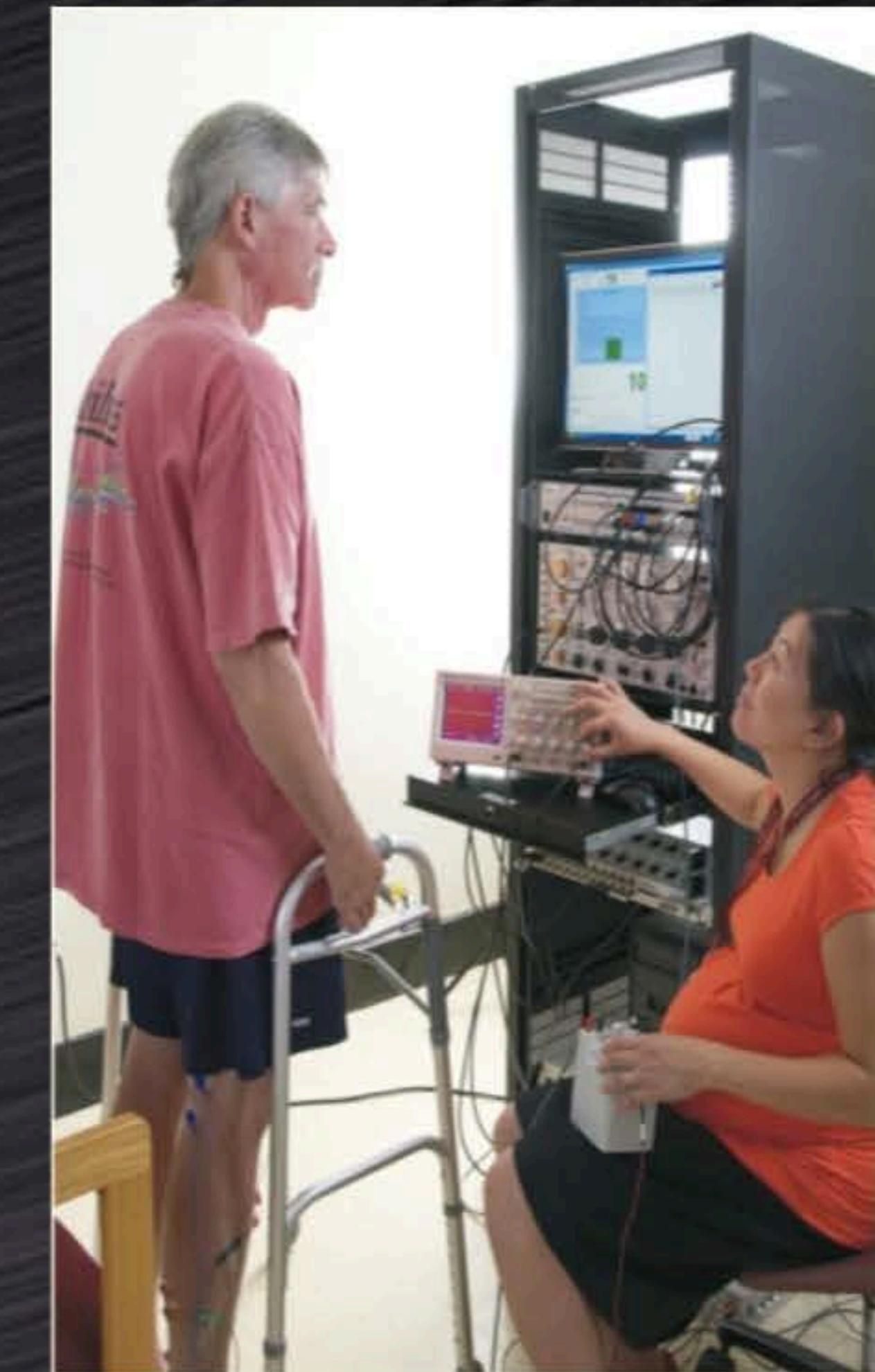
Monkey stretch reflex  
1983



Rat H-reflex  
1995



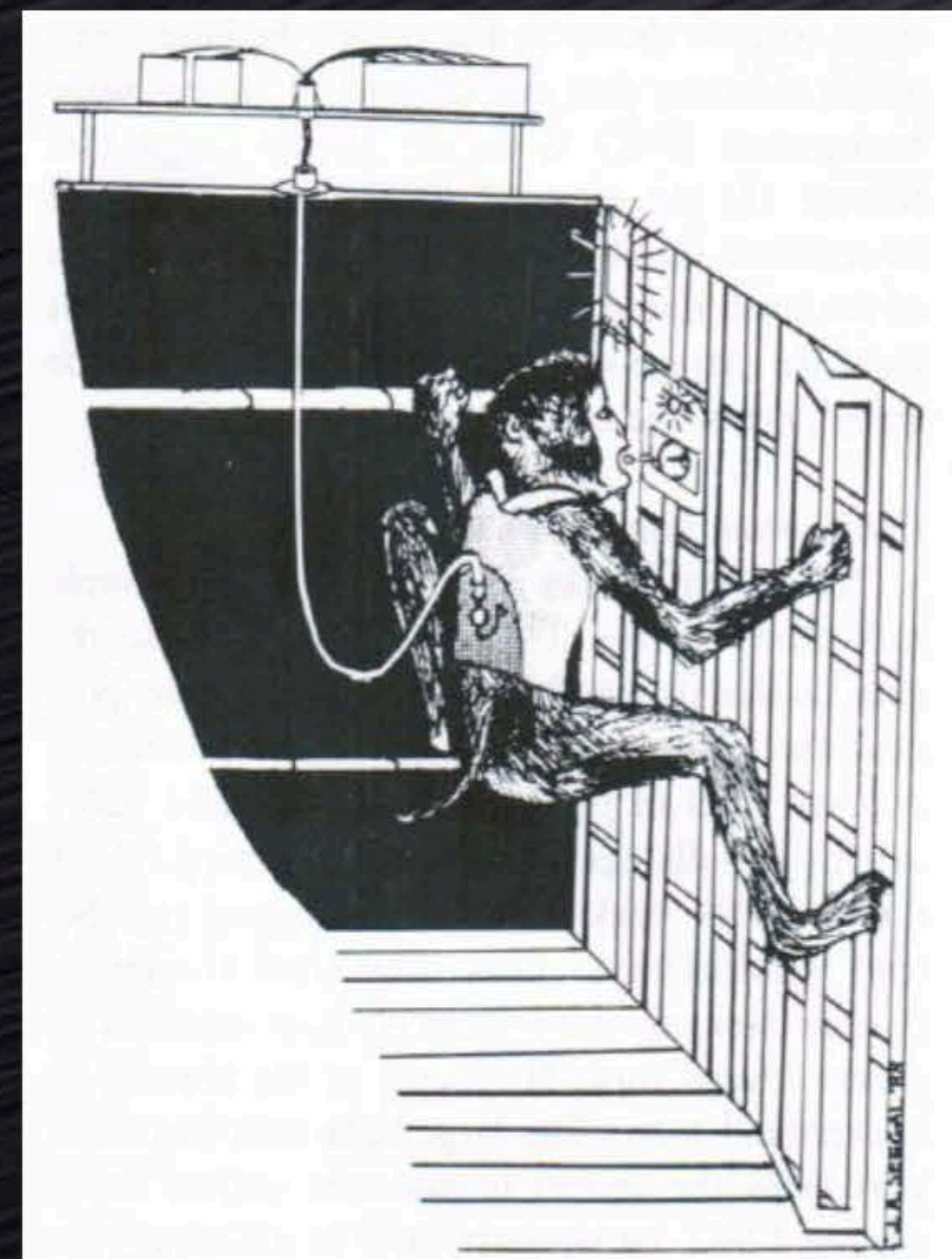
Rats with SCI  
2006



People with SCI  
2013

*MODEL DEVELOPMENT* ———→ *MECHANISTIC STUDIES* ———→ *THERAPEUTIC APPLICATIONS*

1987  
Monkey H-reflex



2005  
Mouse H-reflex



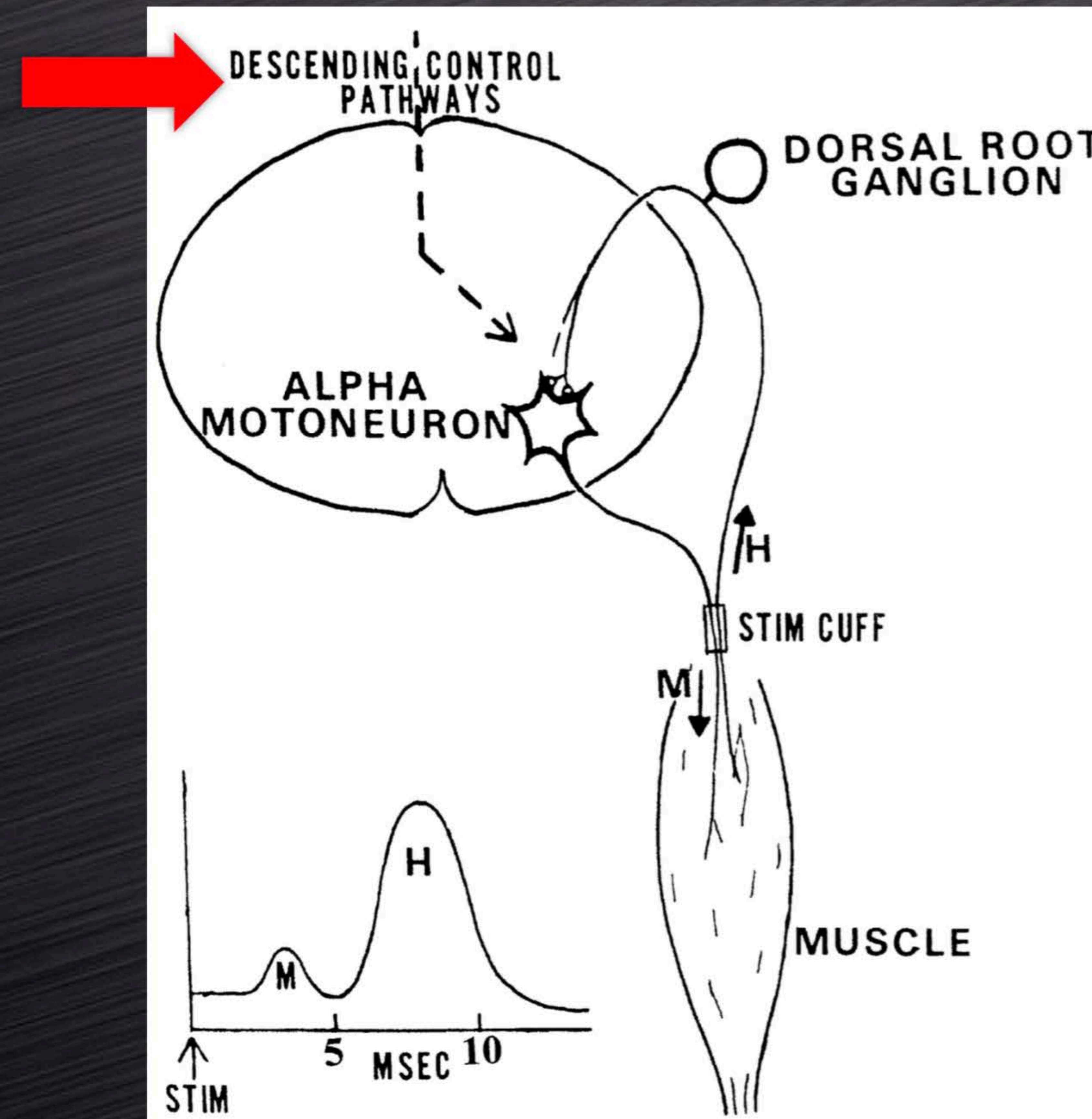
2009  
Human H-reflex





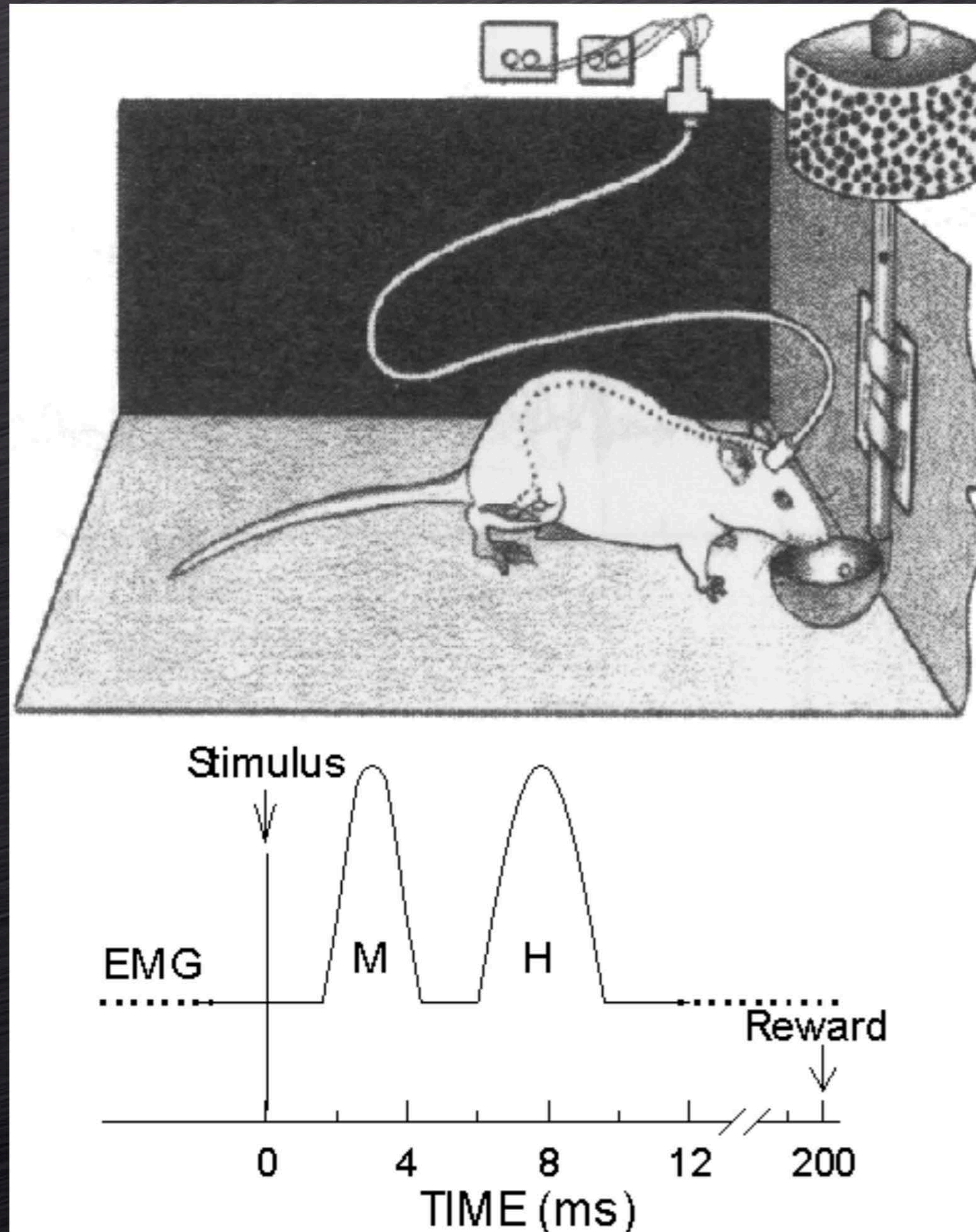
# Operant Conditioning

A subject is given a specific stimulus or placed in a specific situation, and reinforcement (reward) occurs when a specific response occurs. After repeated exposures to this experience, the required response occurs more frequently and thereby increases the number of rewards.





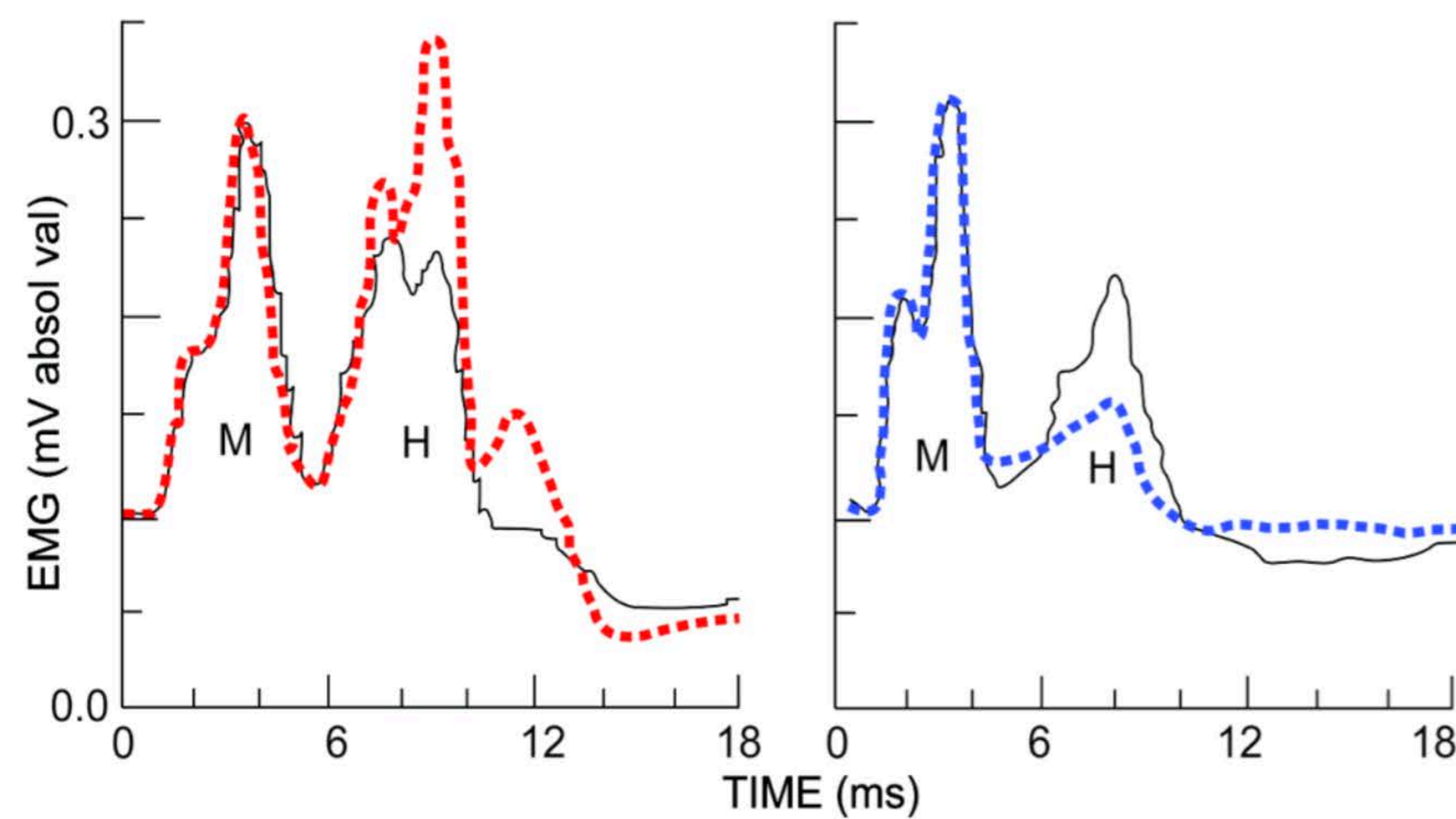
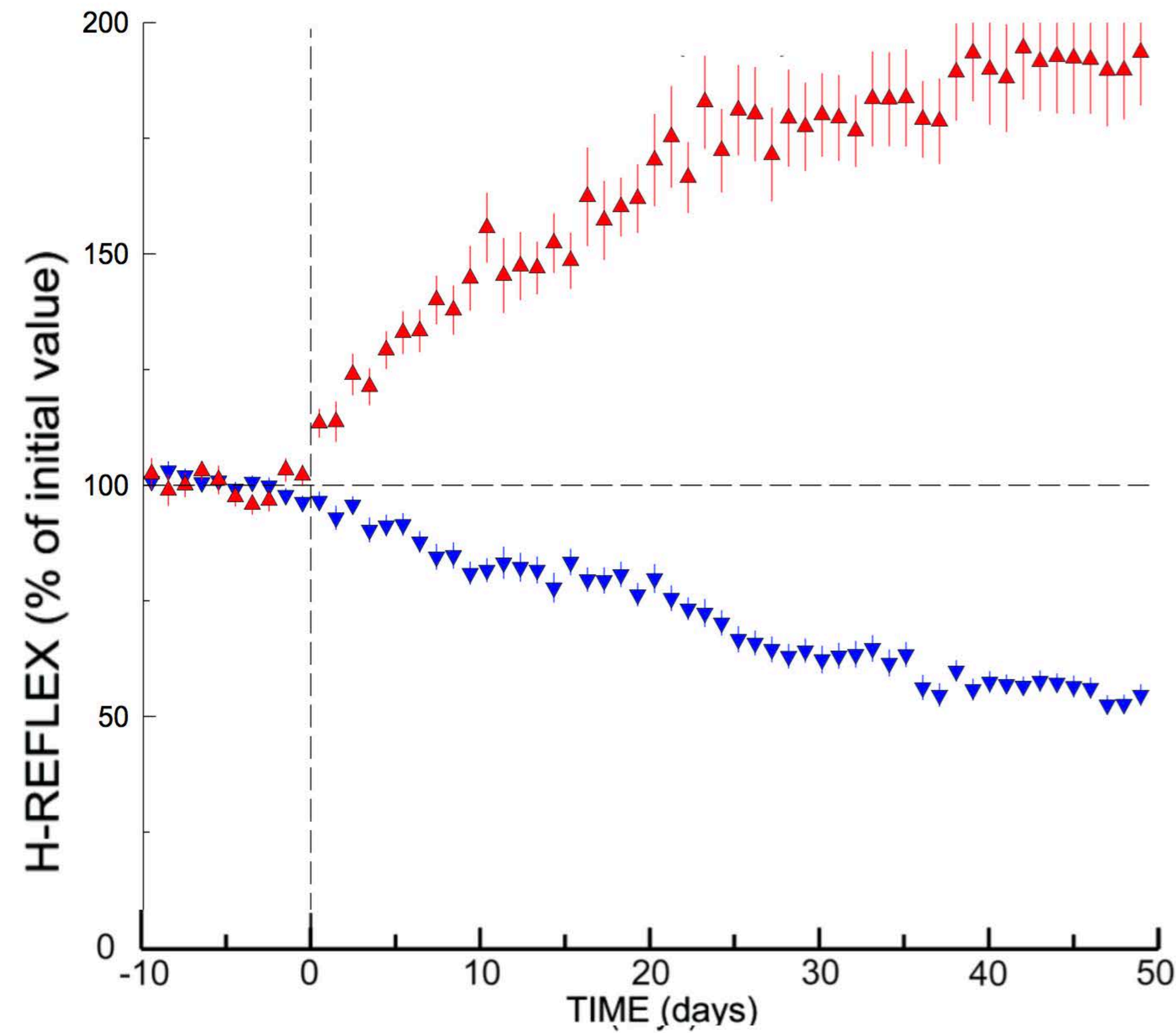
# The Operant Conditioning Protocol



Wolpaw TINS 1997

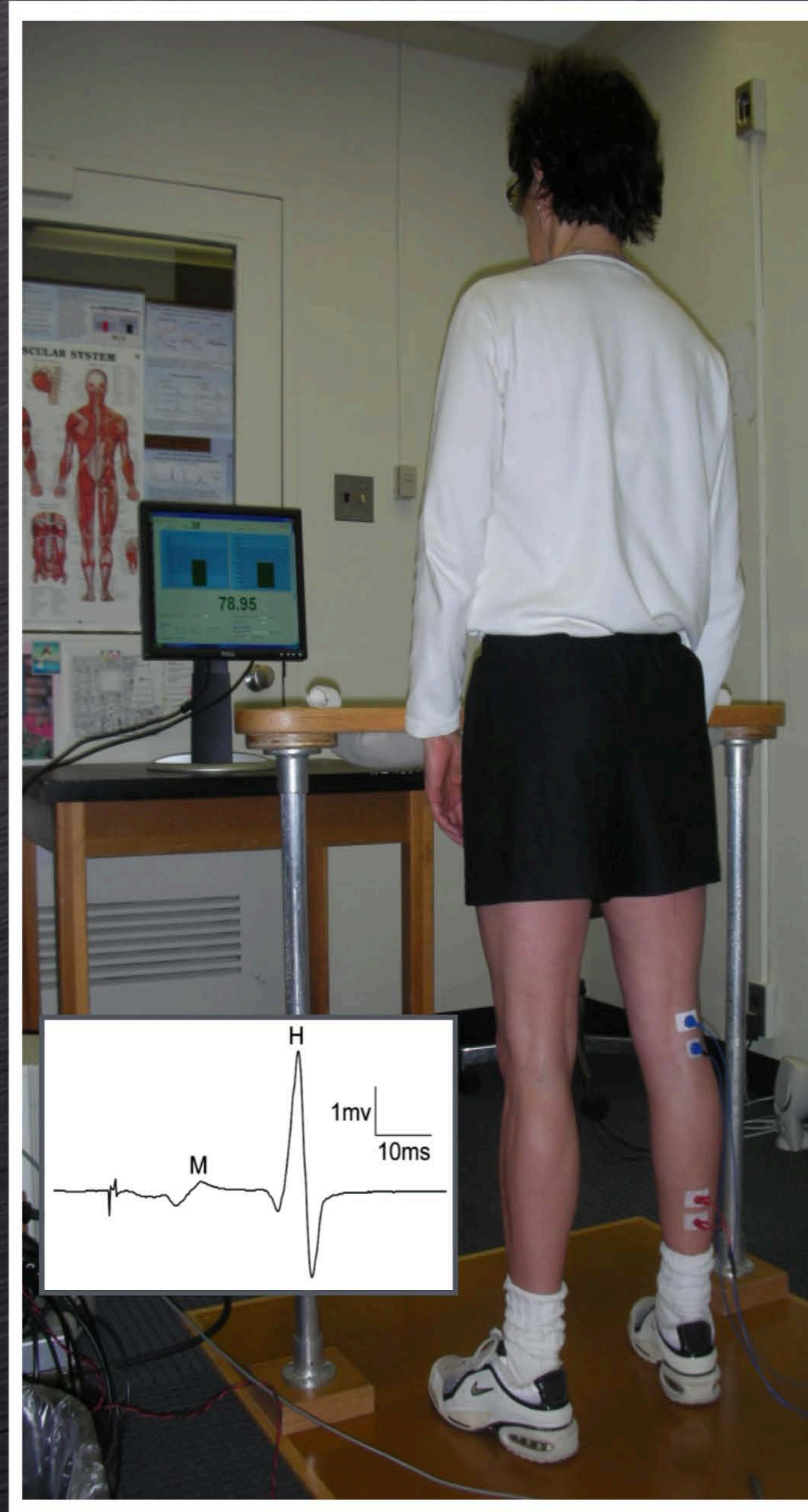


# H-Reflex Conditioning in Rats





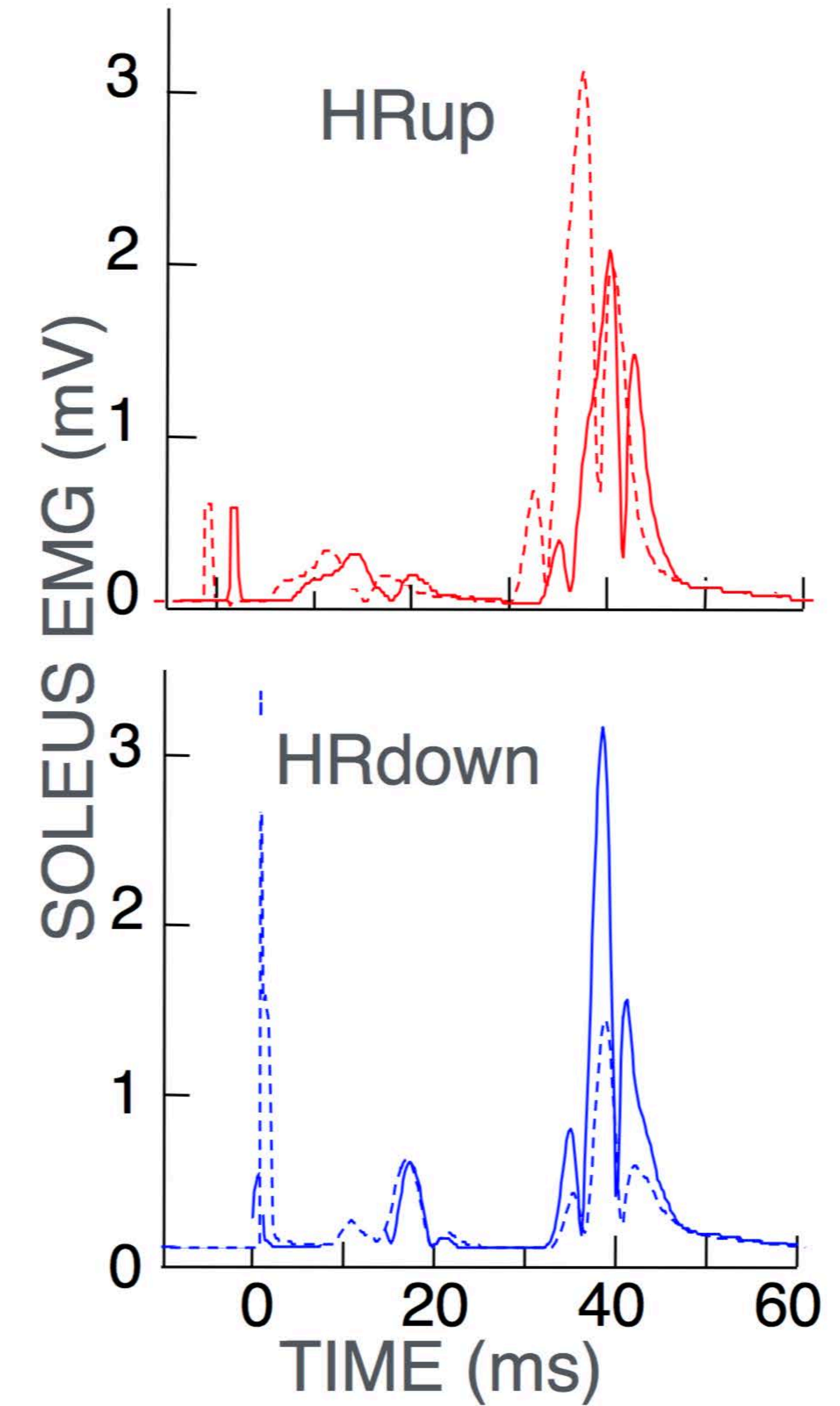
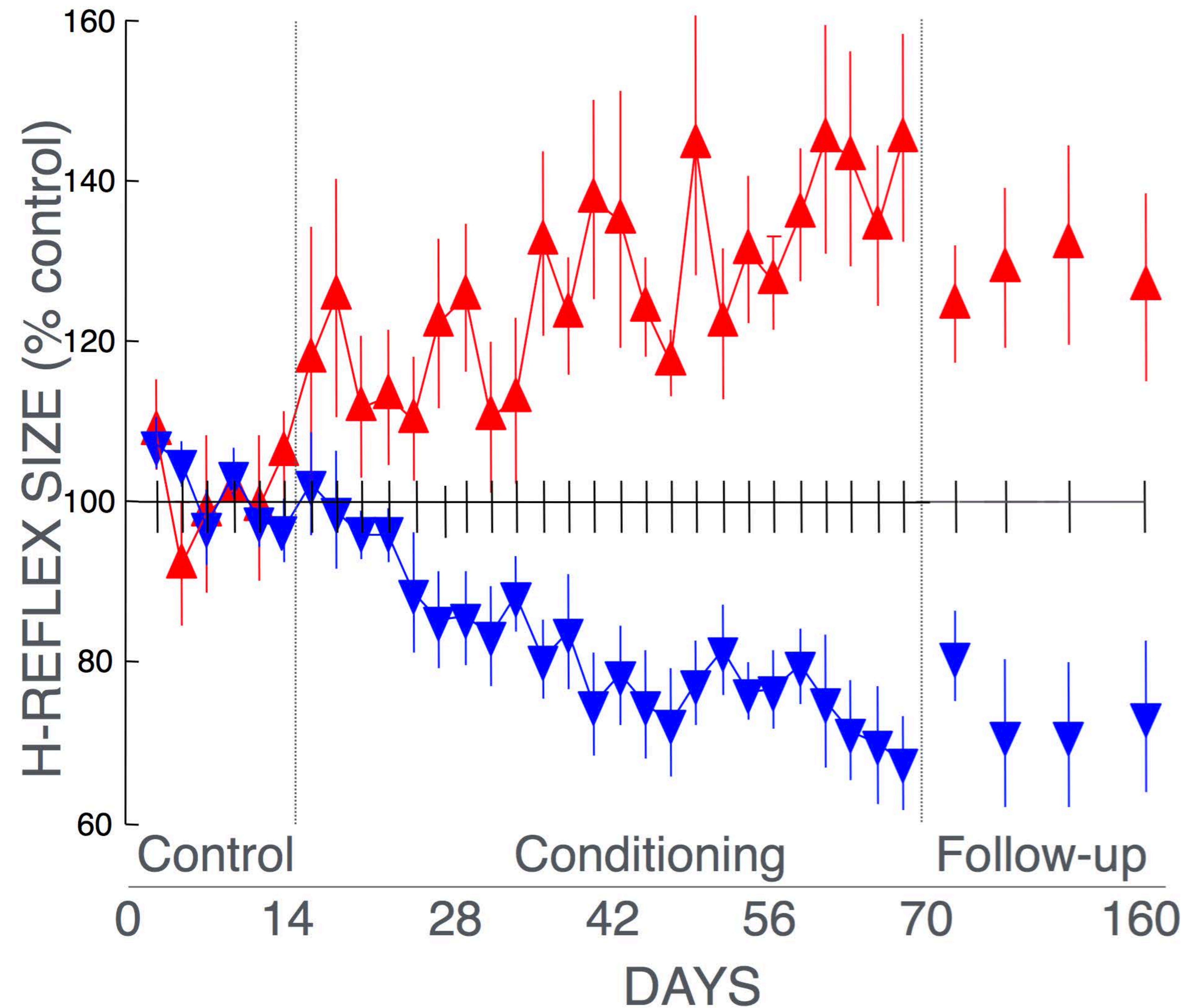
# H-Reflex Conditioning in People



Thompson et al. J Neurosci 2009

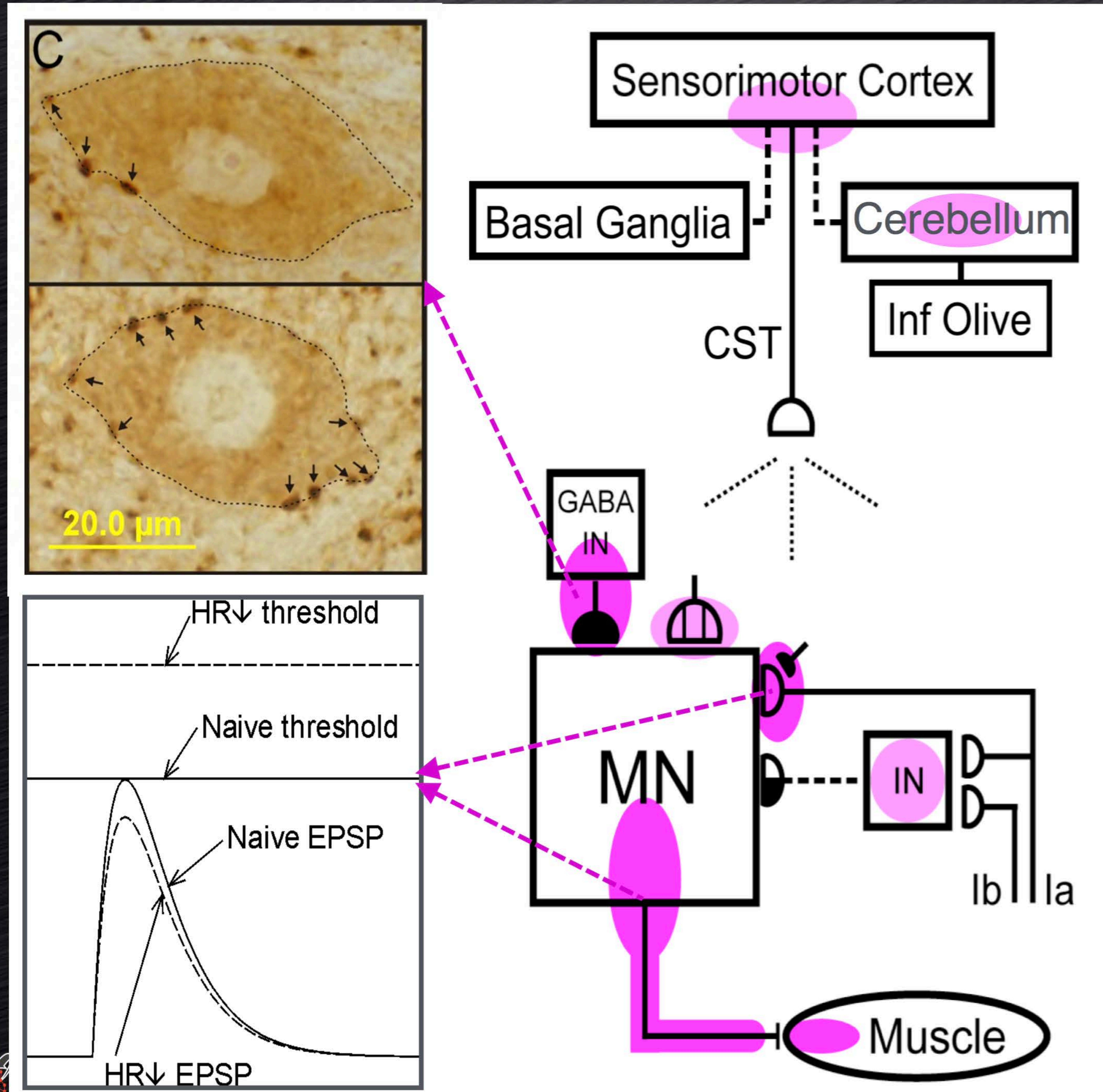


# H-Reflex Conditioning in People





# H-reflex conditioning creates a hierarchy of plasticity

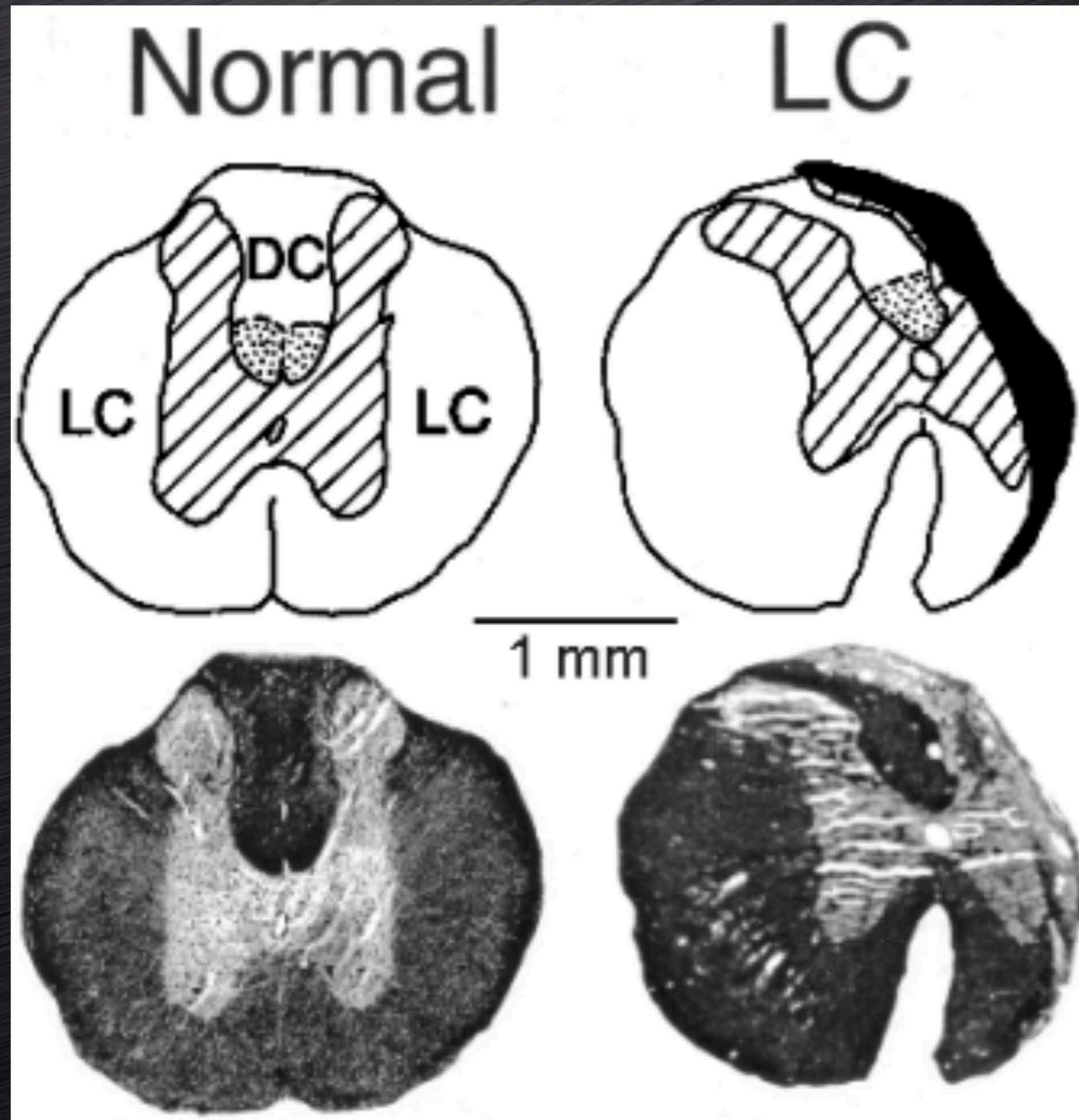


Updated from  
Encyclopedia of  
Neuroscience, 2009





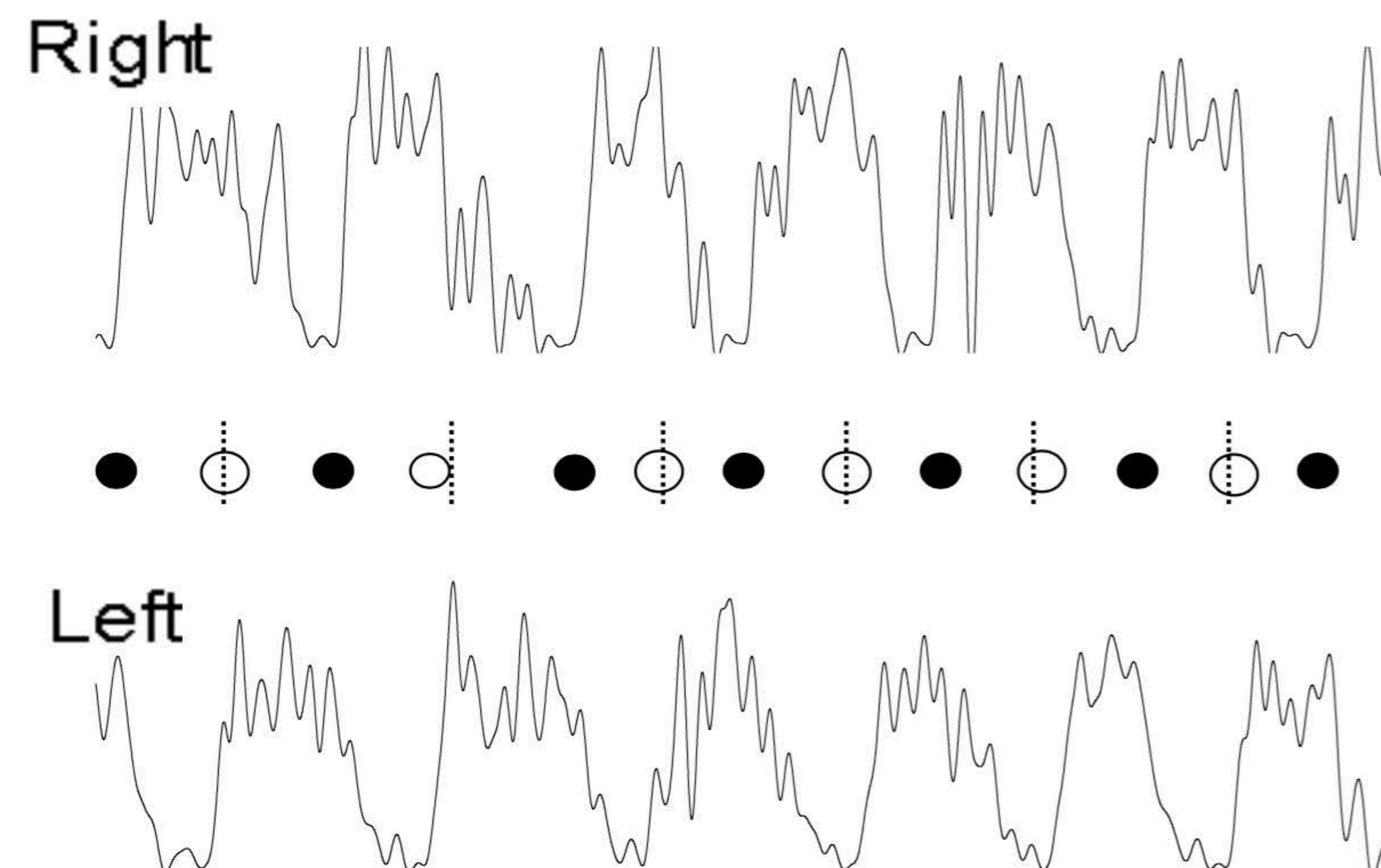
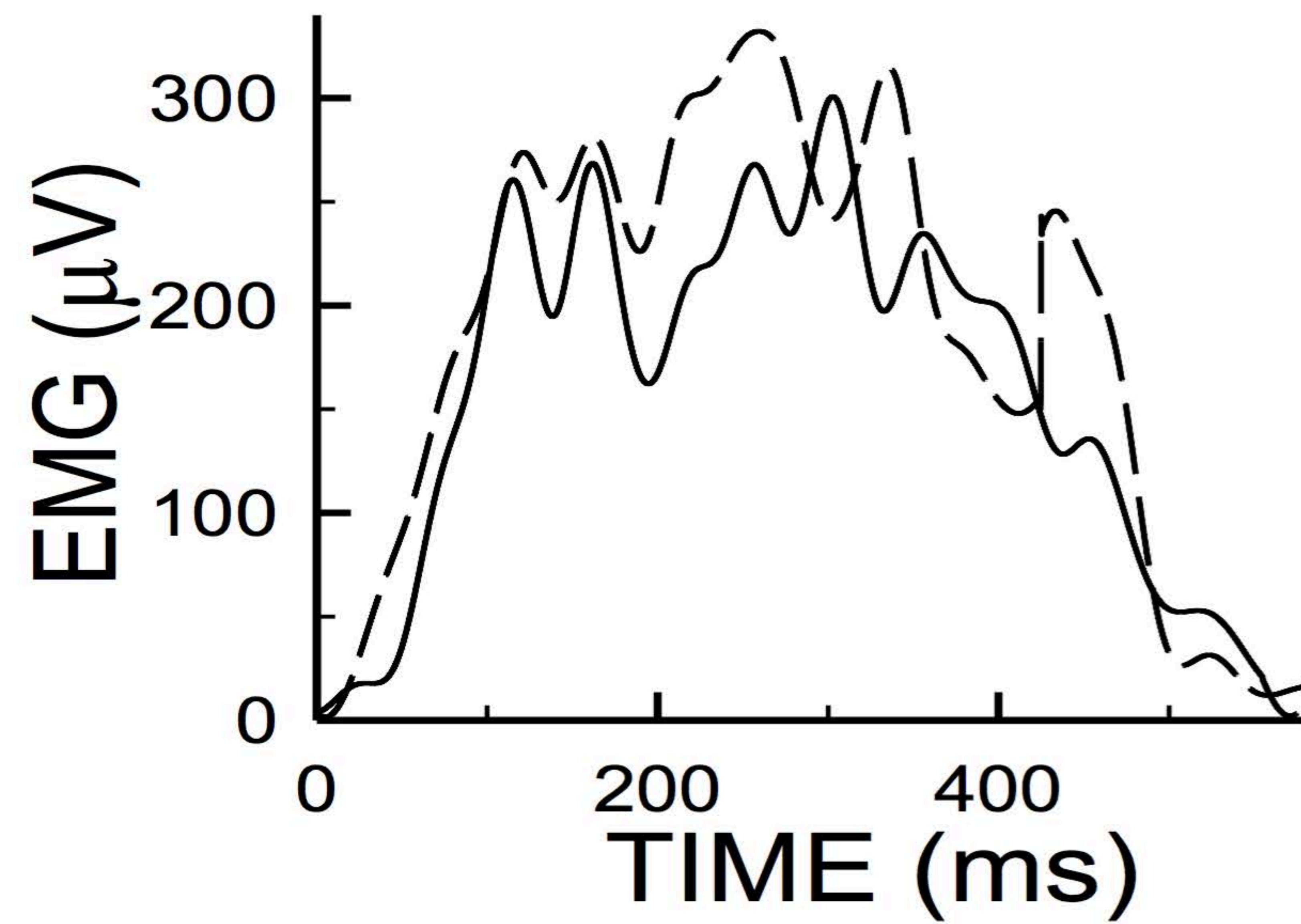
# Spinal Cord Injury: T9-10 Lateral Column Transection



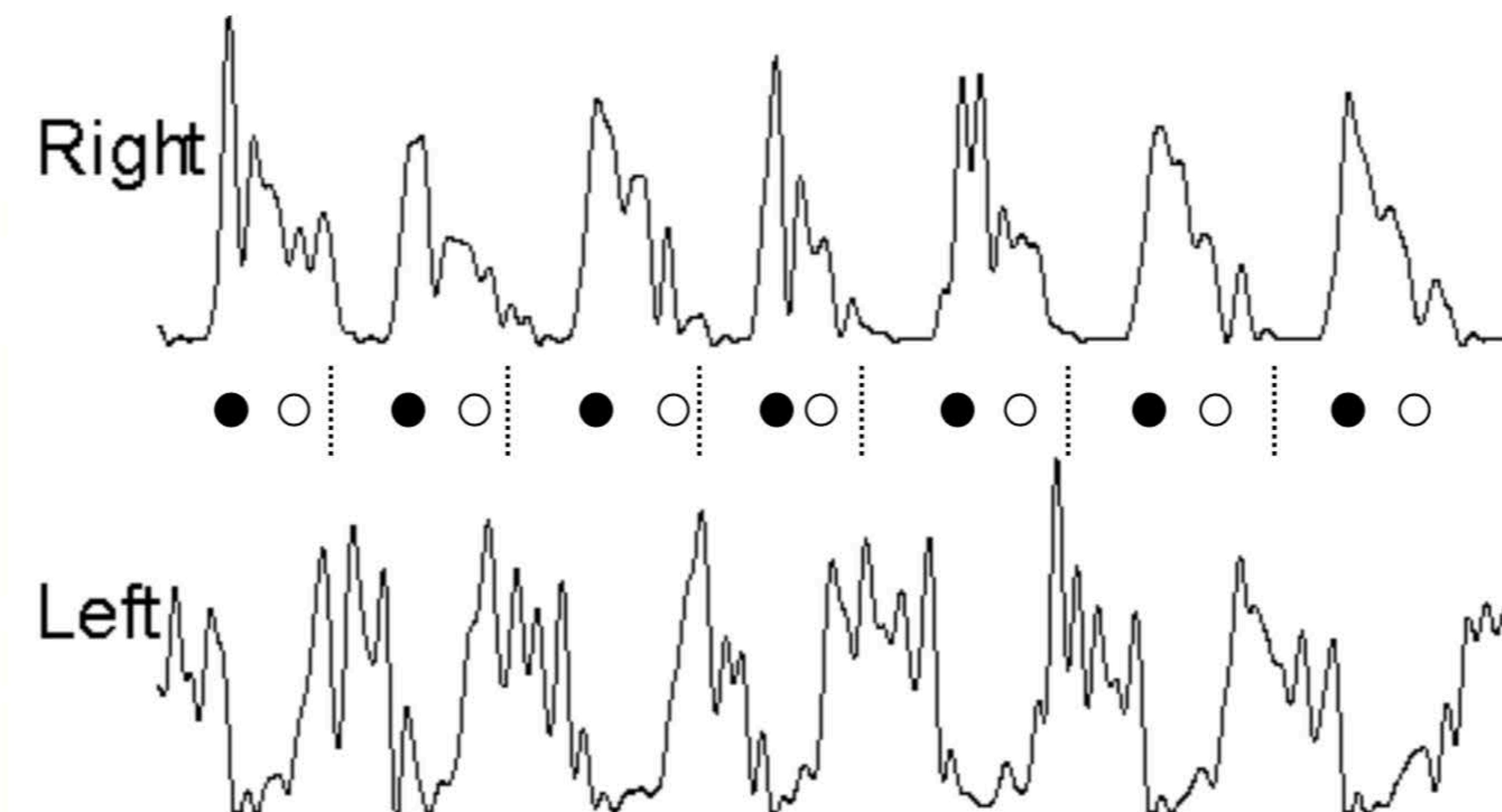
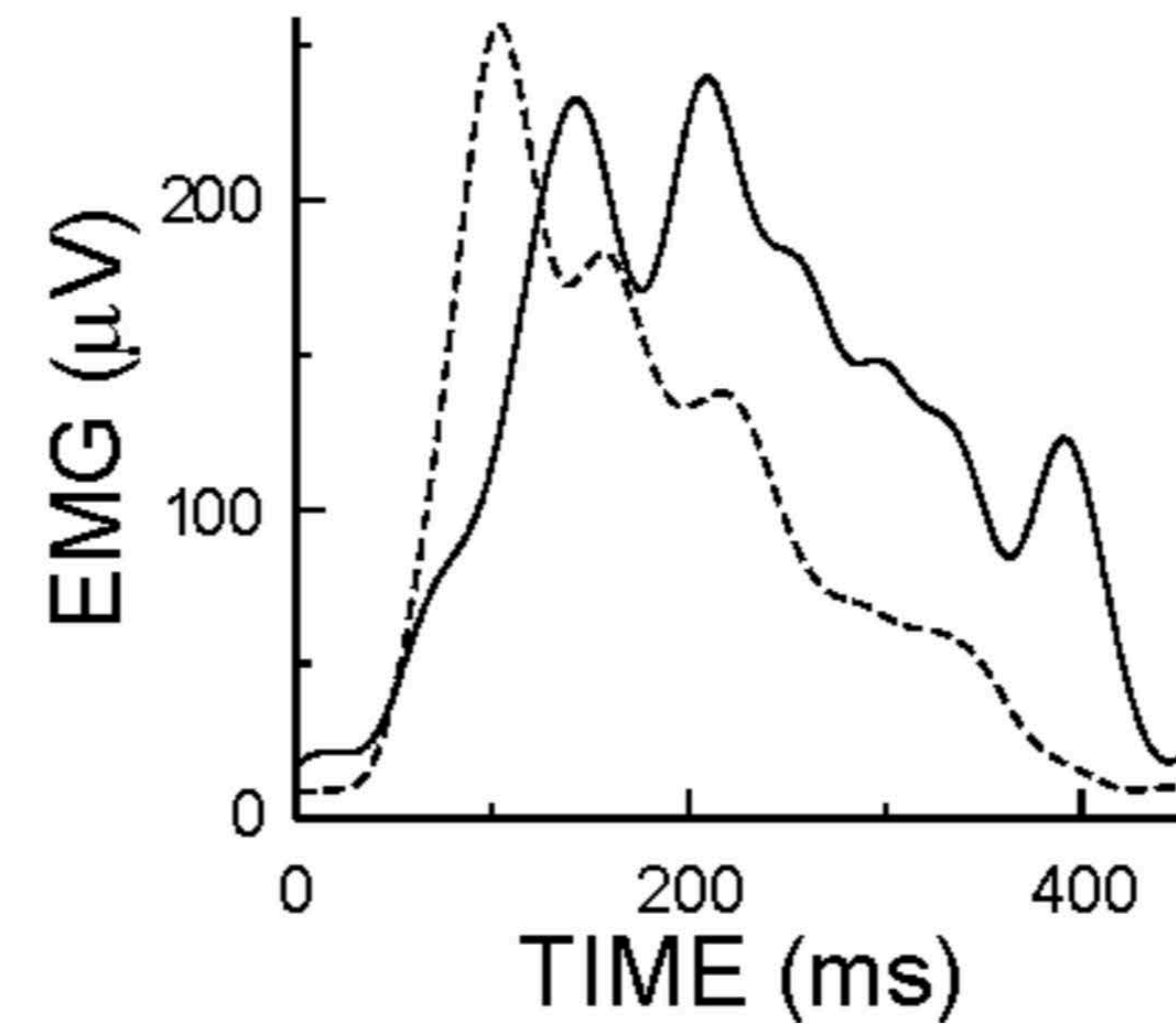
Chen et al J Neurosci 2006



## Normal Rat

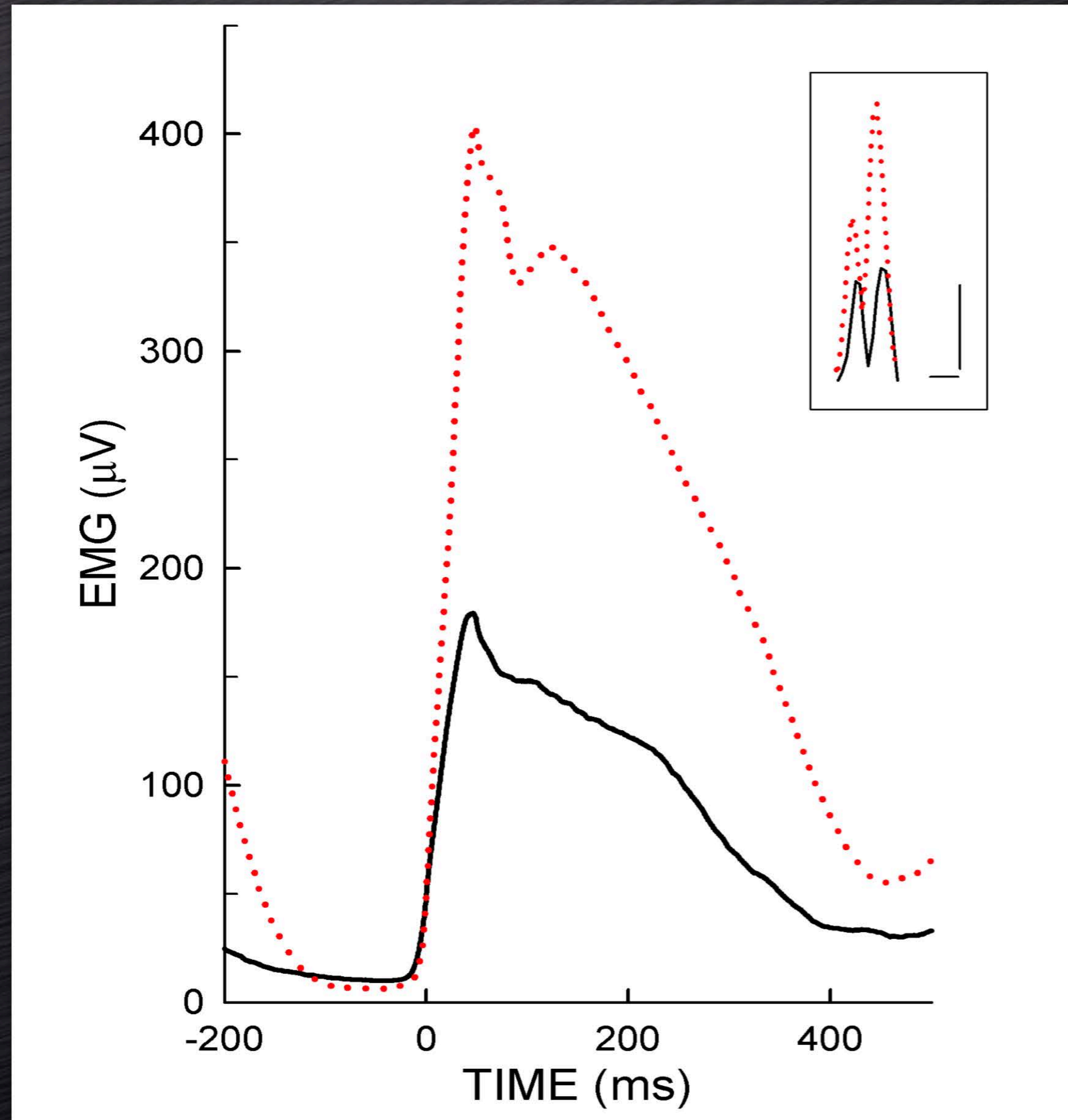


## Rat with Right LC Transection



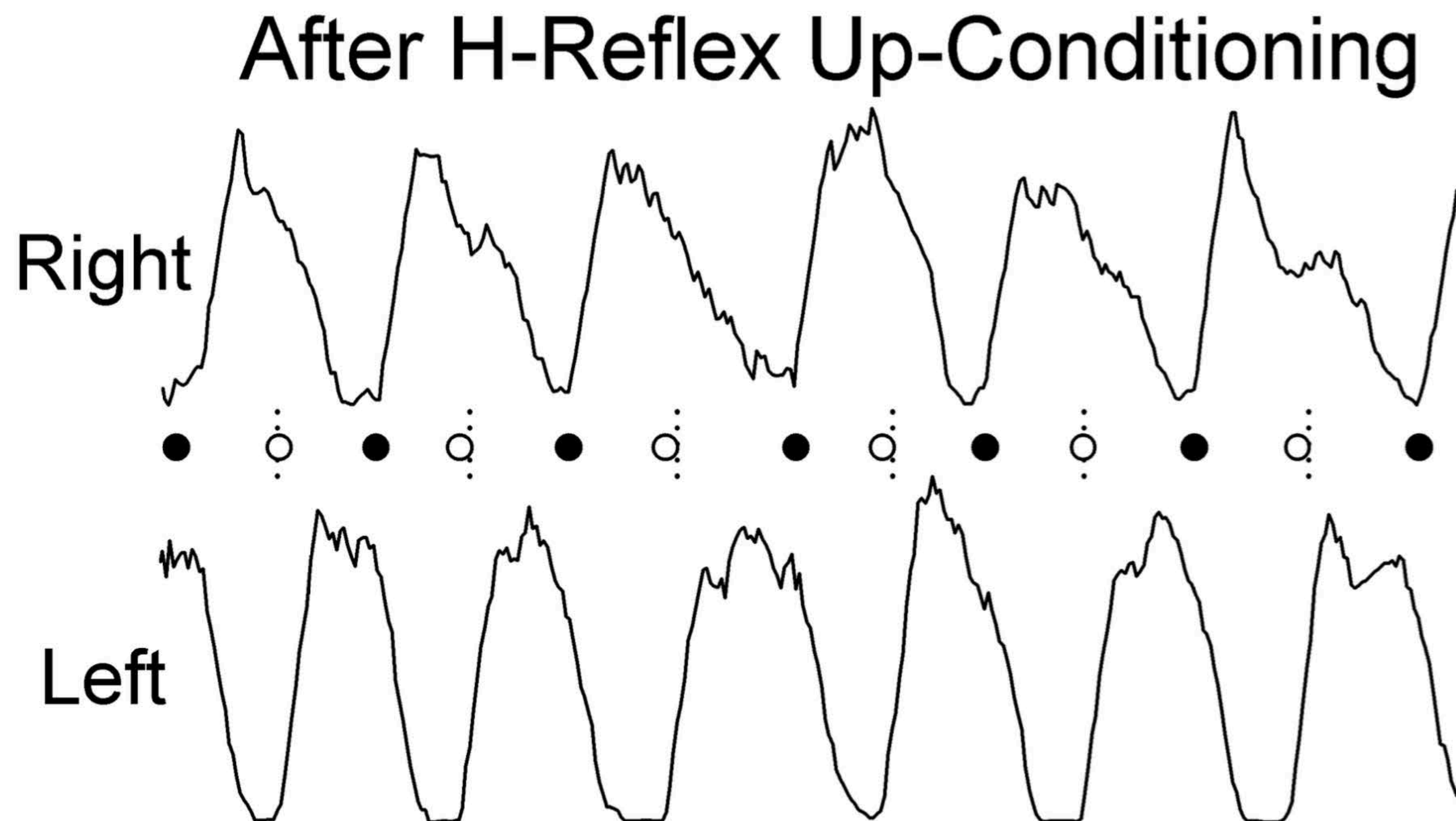
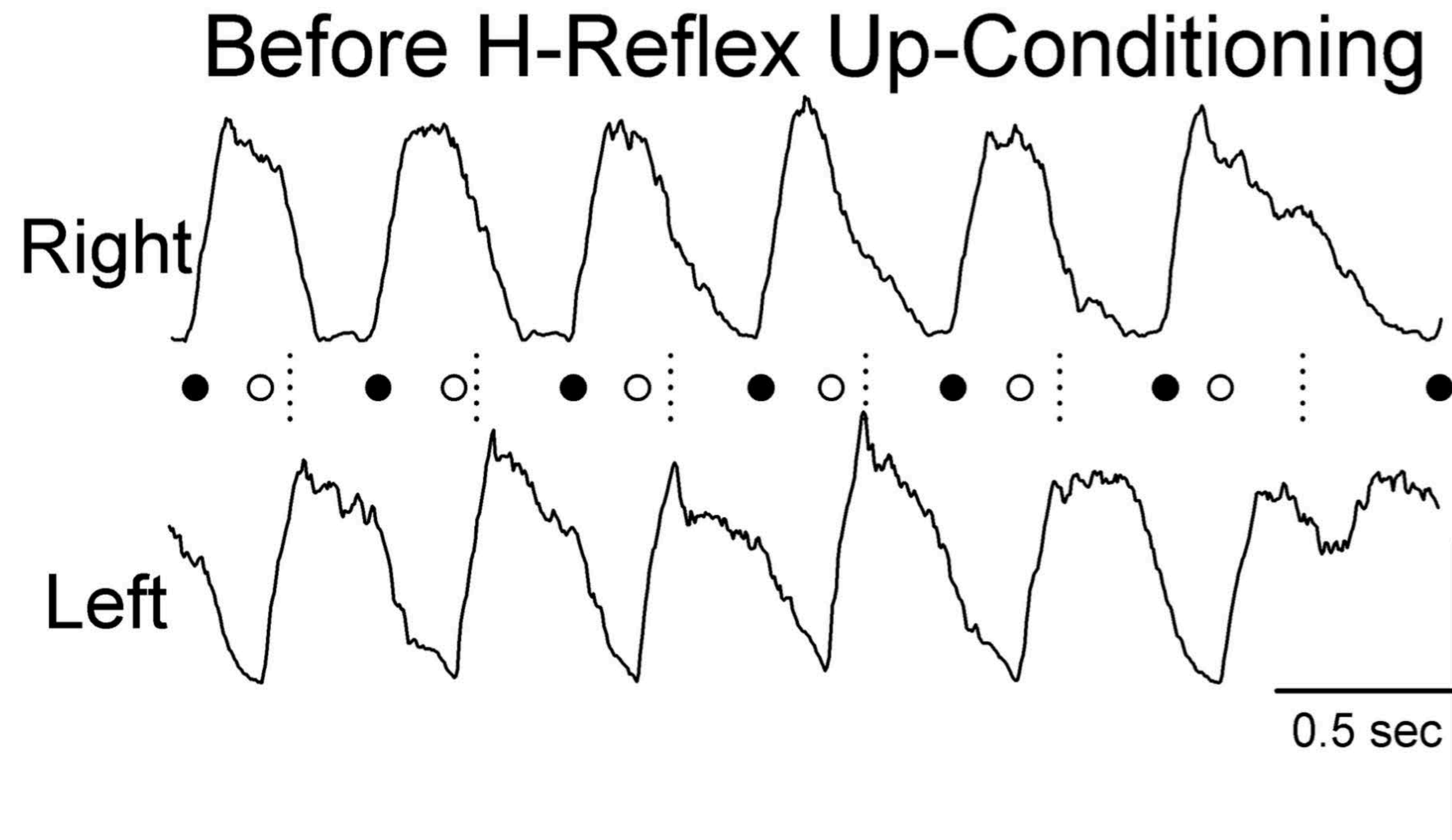


# HR up-conditioning increases the soleus burst in an LC rat





# Up-Conditioning in an LC Rat

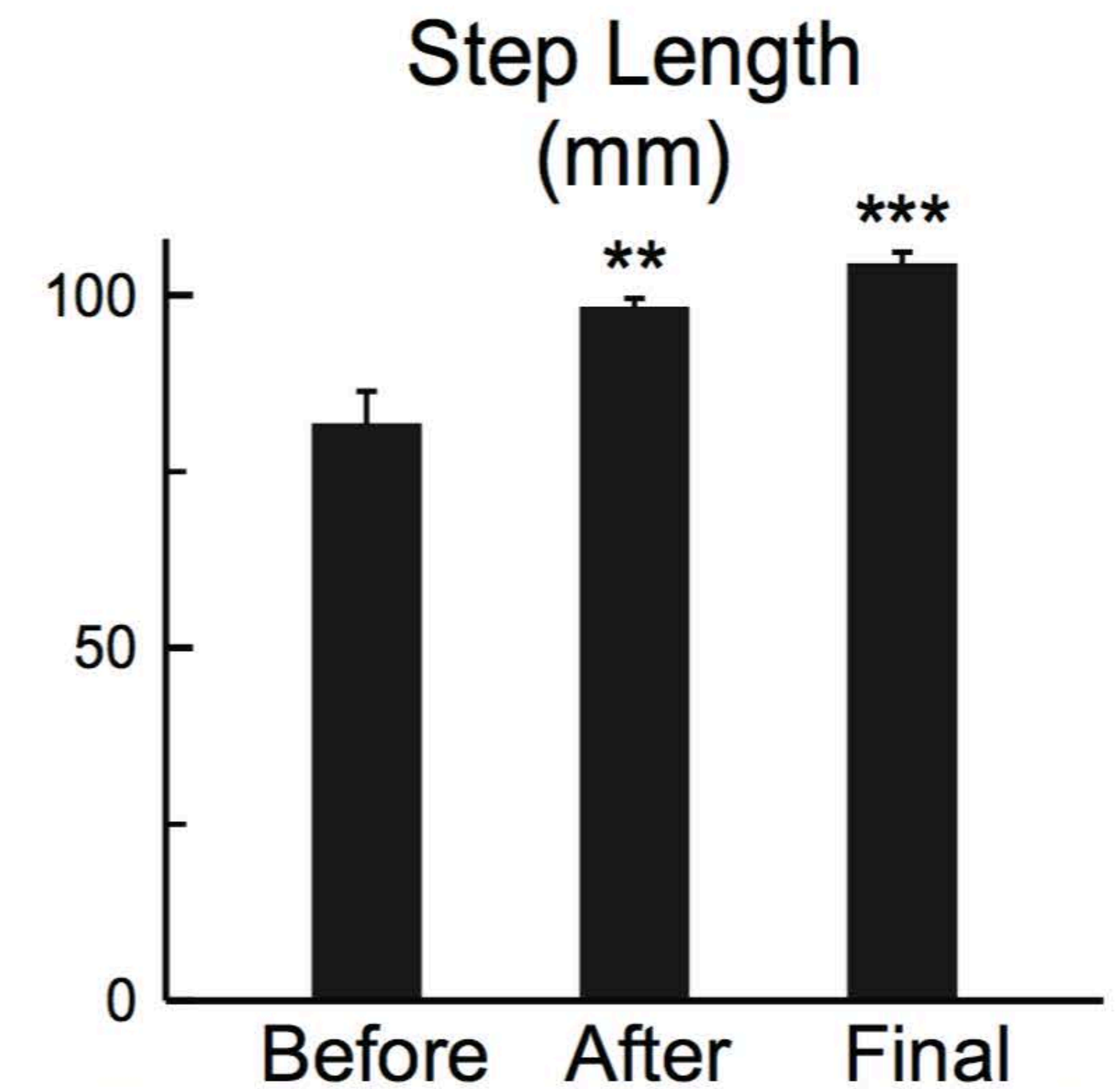
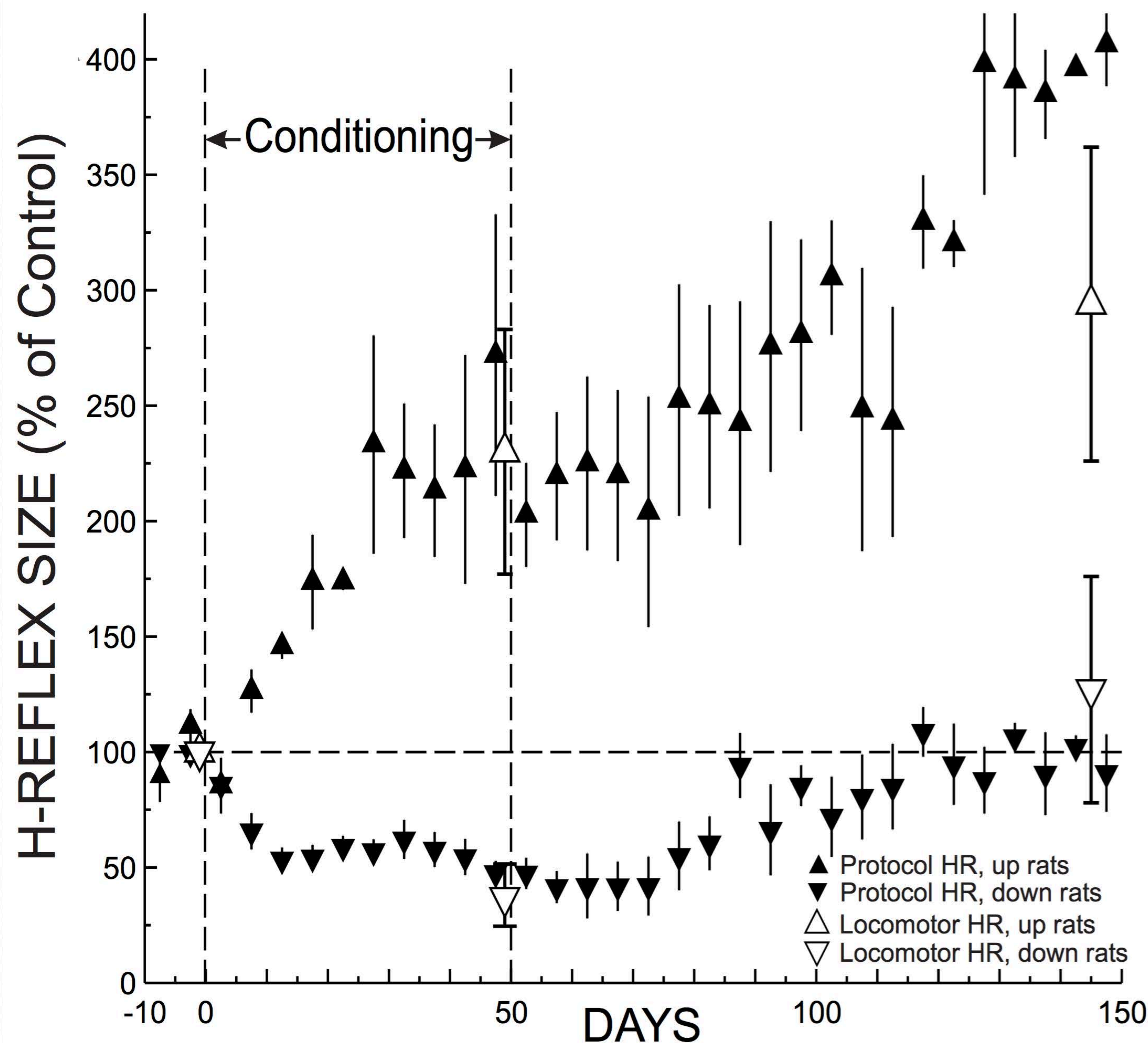


Appropriate reflex conditioning restores gait symmetry.

Chen et al,  
J Neurosci 2006



After conditioning ends, beneficial effects persist & increase.



Chen et al,  
J Neurophysiol 2014



# *What about Humans?*

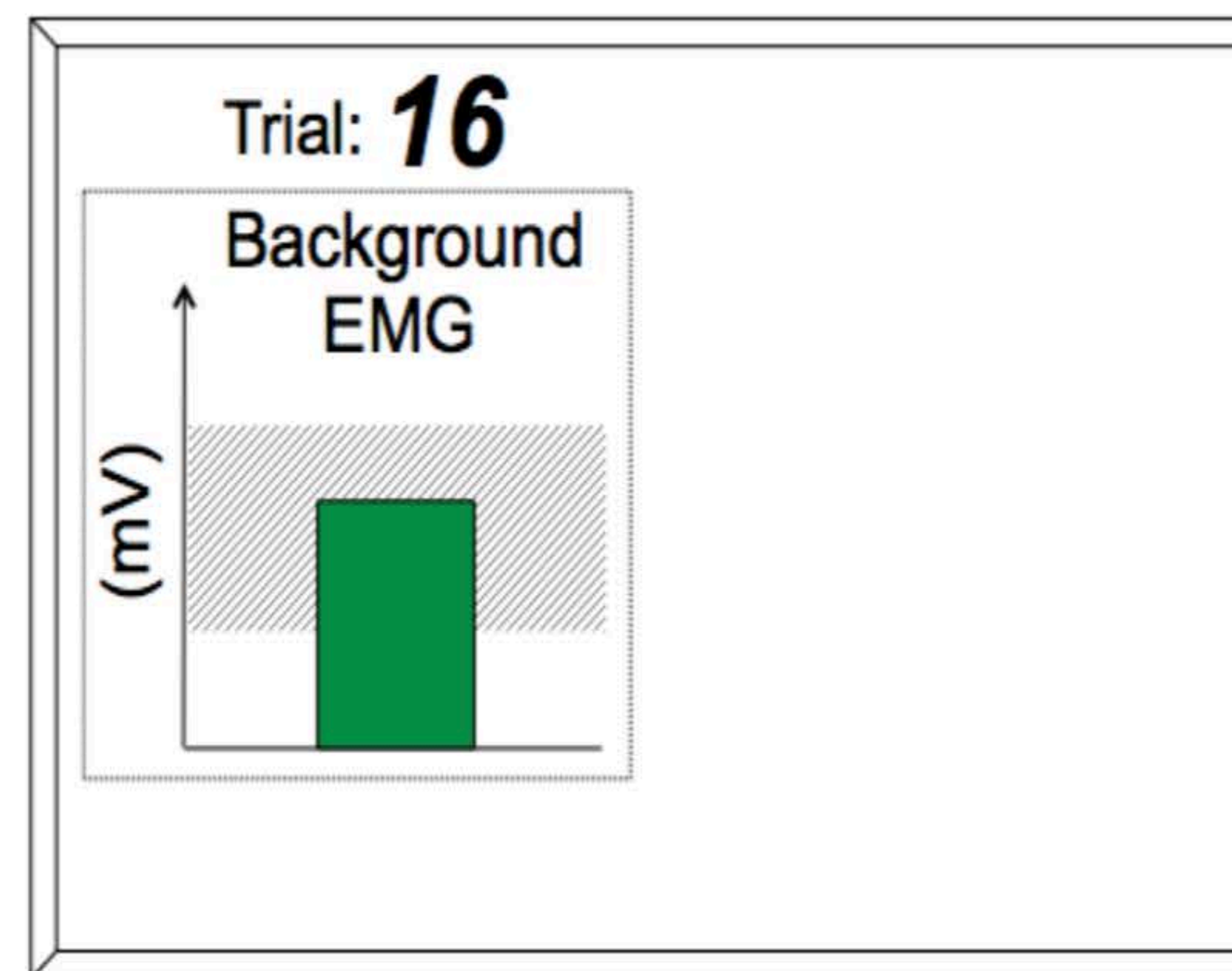
*In people with spasticity and footdrop due to SCI,  
can soleus HRdown conditioning improve locomotion?*



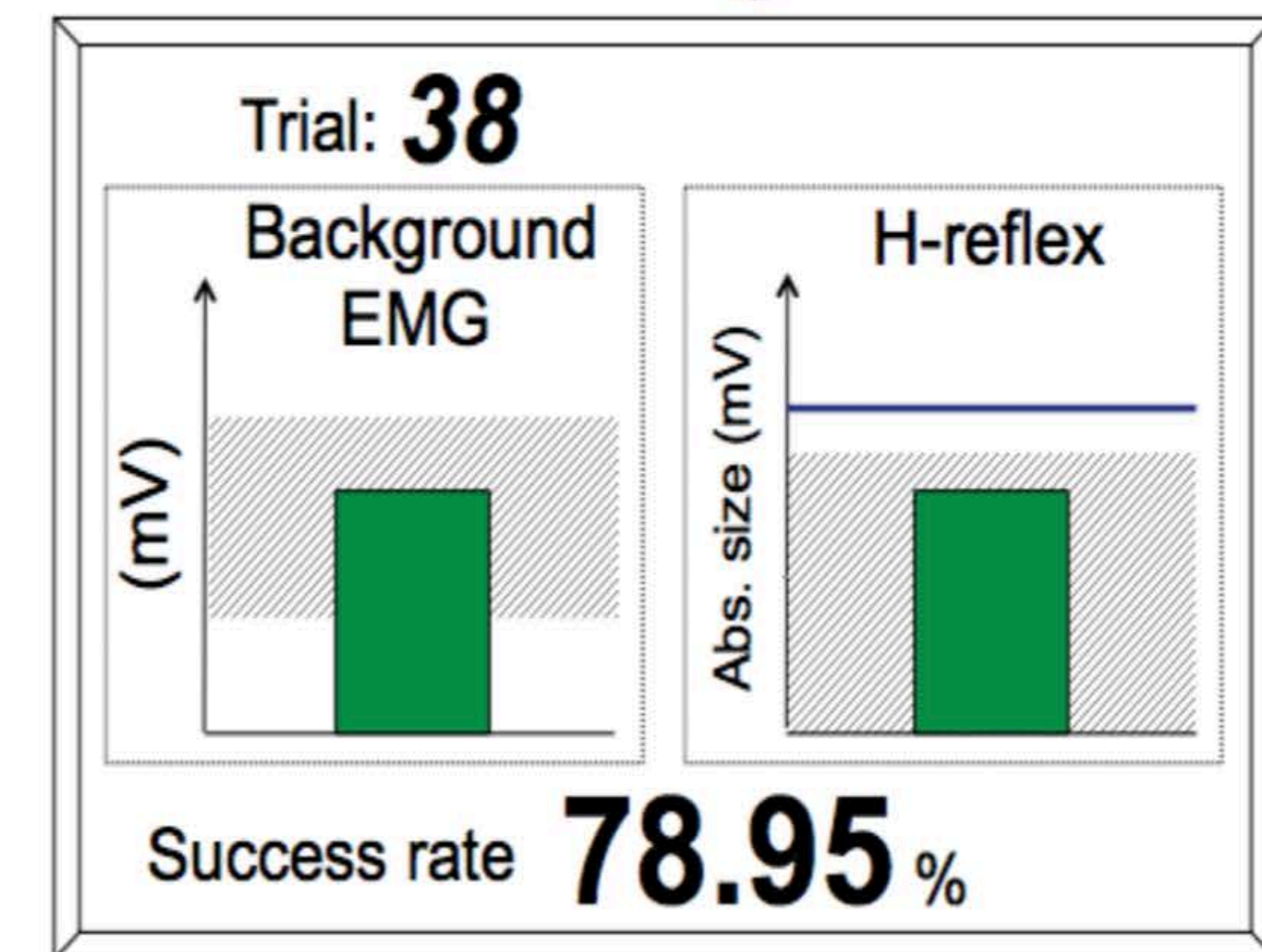
# Format



## Control trials



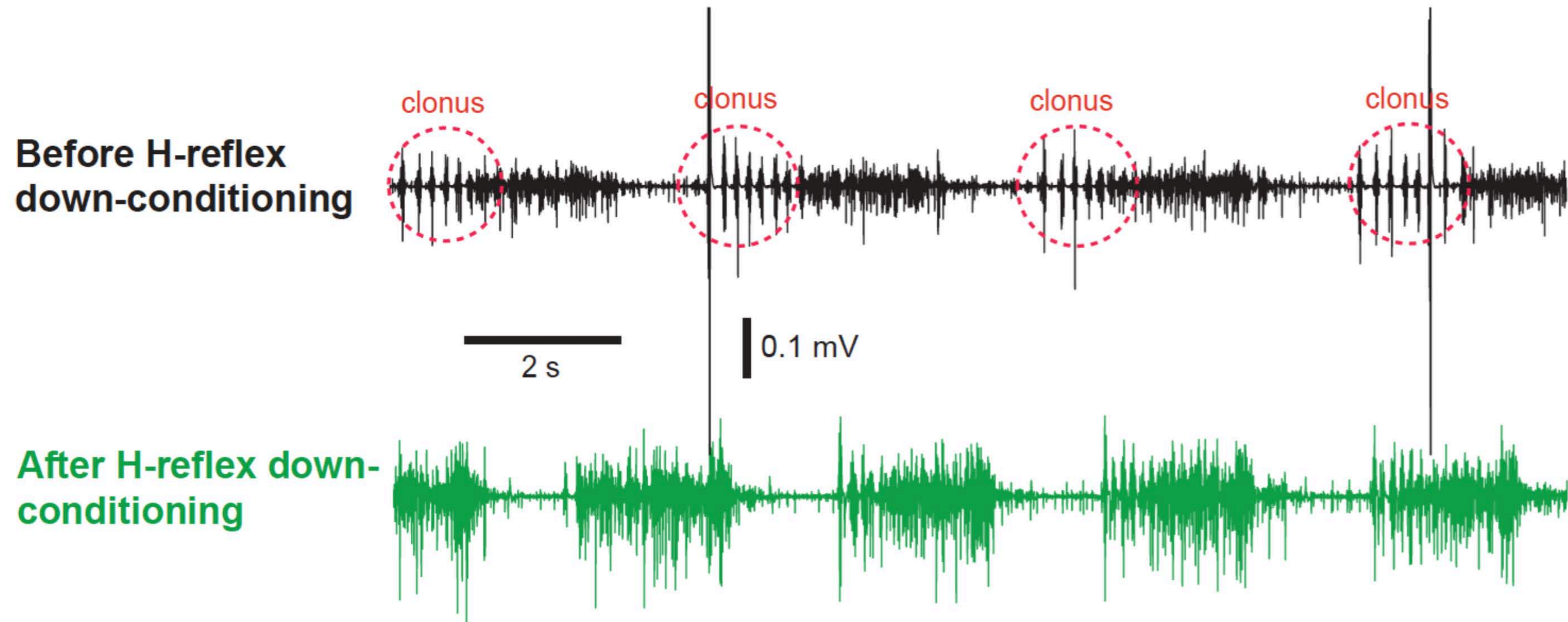
## Conditioning trials



Thompson et al J Neurosci 2013

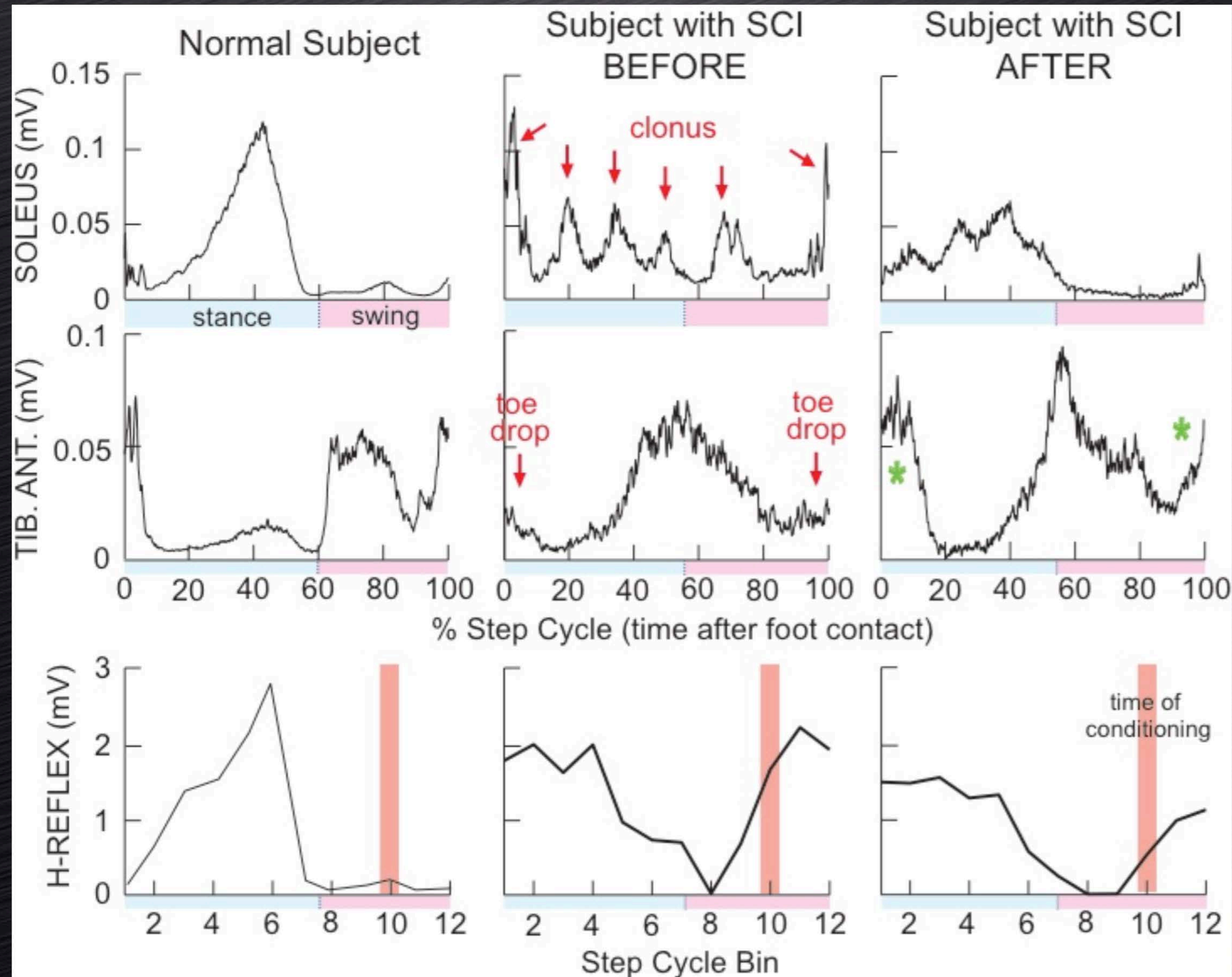


# Improved soleus locomotor EMG activity



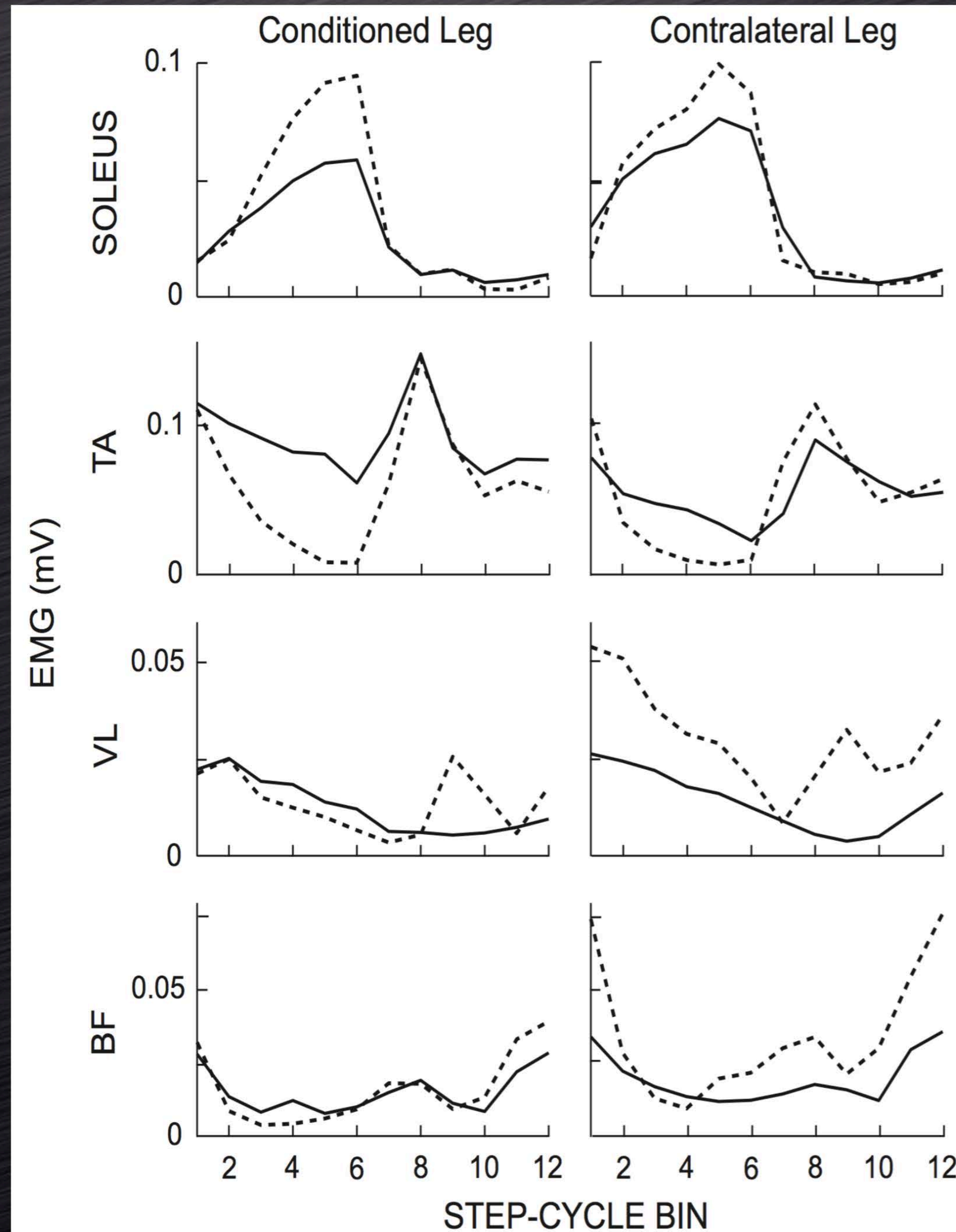


# Improved soleus and tibialis anterior EMG after soleus H-reflex down-conditioning





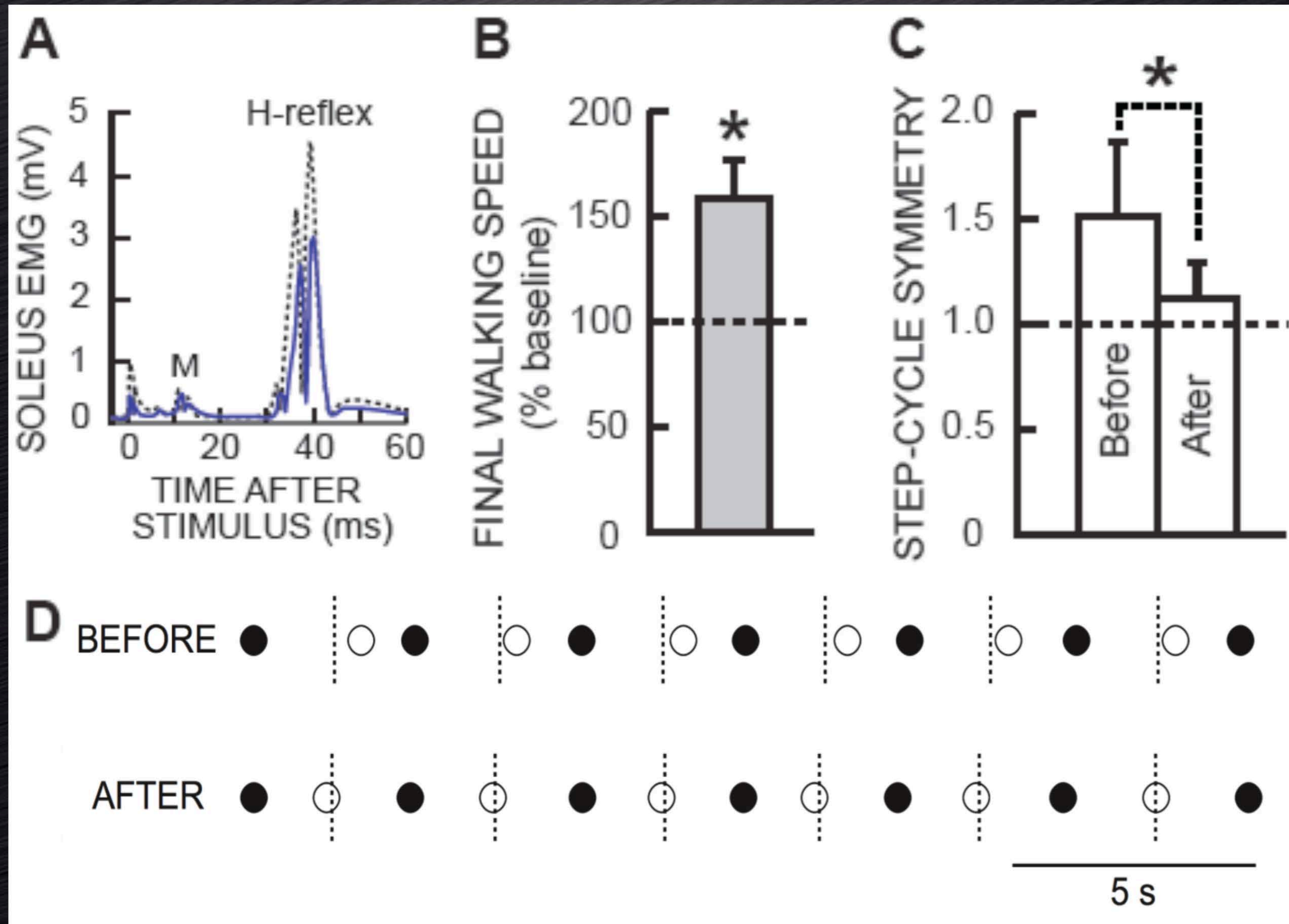
# The targeted plasticity triggers widespread plasticity



Thompson et al  
J Neurosci 2013

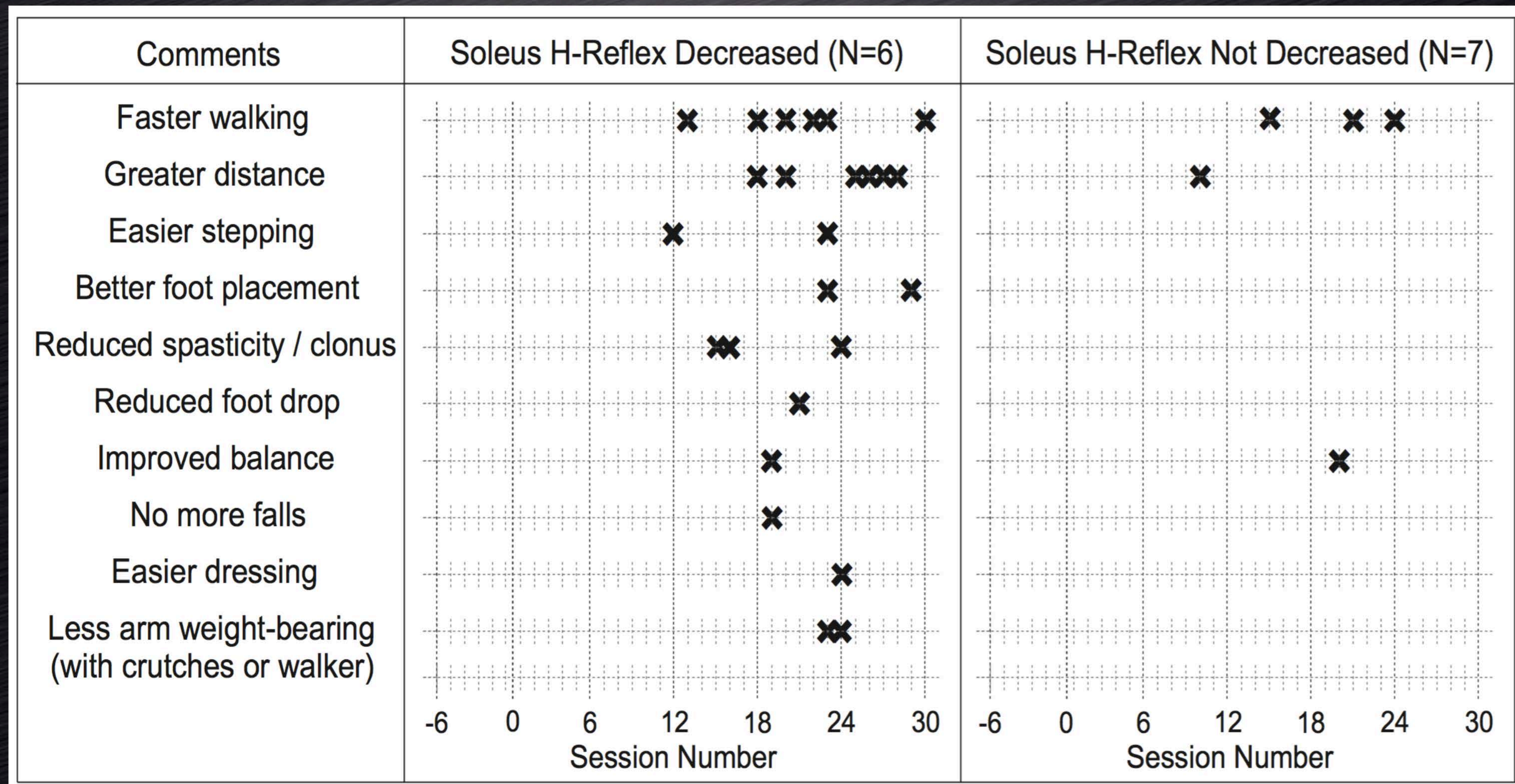


# Walking speed & symmetry improve





# Spontaneous Comments



*Gait improvement is apparent to the subjects.*





Thompson et al.  
Helen Hayes / Wadsworth Center



# Summary

- Recent scientific & technical advances enable a new class of technologies that interact directly with the CNS.
- Brain-computer interfaces (BCIs) to restore communication & control.
- BCIs to enhance rehabilitation.
- Safe rapid cortical mapping prior to surgery.
- Protocols that target beneficial plasticity to critical sites and trigger wider beneficial plasticity.



# CREDITS



Aiko  
Thompson  
Med Univ SC



Bill Haug  
Med Univ SC



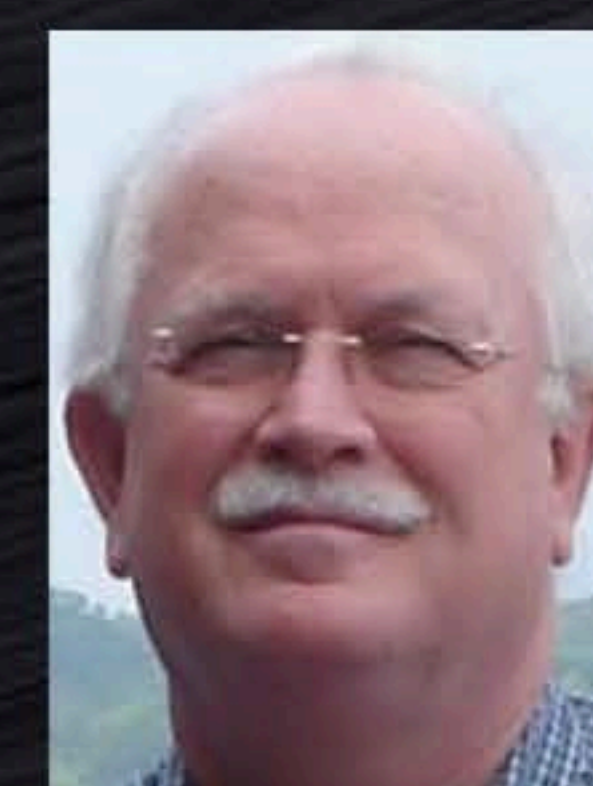
Dave  
Reinkensmeyer  
UC Irvine



Sumner Lee  
Norman  
UC Irvine



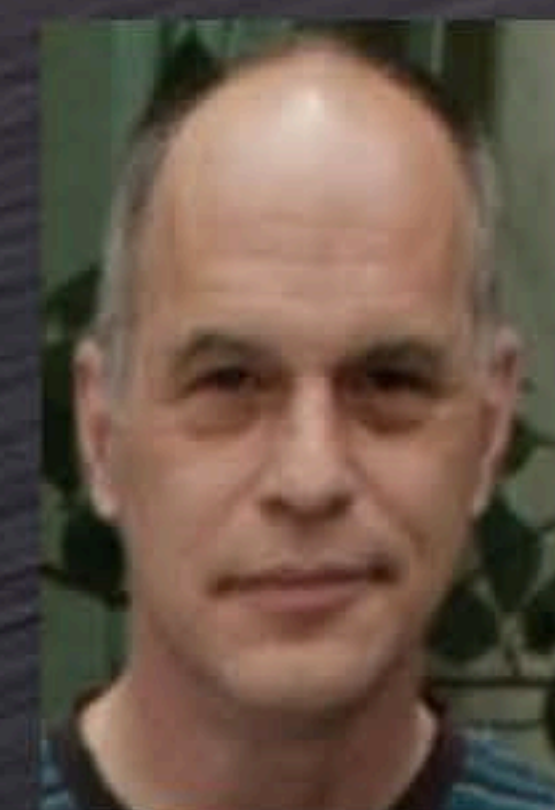
Brad Voytek  
UCSD



Art English  
Emory Univ



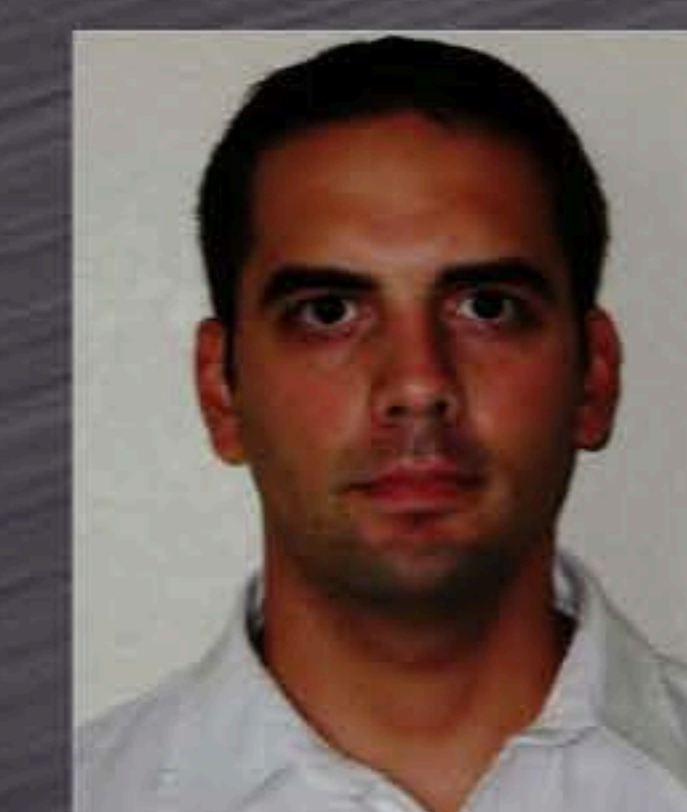
Jason Carmel  
Cornell Univ



Ayse Gunduz  
Univ Fl  
Gainesville



Brian Litt  
Univ Penn



Dean Krusienski  
Old Dominion  
Univ



Adrianne Randolph  
Kennesaw State Univ



Rita Goldstein  
Mount Sinai



Mohammad  
Parvaz  
Mount Sinai



Eric  
Wolbrecht  
U Idaho



Bob Knight  
UC Berkeley



Jackie Montes  
Columbia Univ



NCAN



# *Recent and Current Support*

NIBIB P41 BTRC Center

NINDS; NCMRR/NICHHD

Department of Veterans Affairs

Army Research Office

DARPA

ALS Hope Foundation

James S. McDonnell Foundation

Altran Foundation

NEC Foundation

Brain Communication Foundation

NYS Spinal Cord Injury Research Trust



# **NCAN**

## **ANNOUNCEMENT**

NIH Short Course in Adaptive Neurotechnologies  
Albany, New York

July 10-28, 2017  
&  
July, 2018



*Thank you!*

*Questions?*